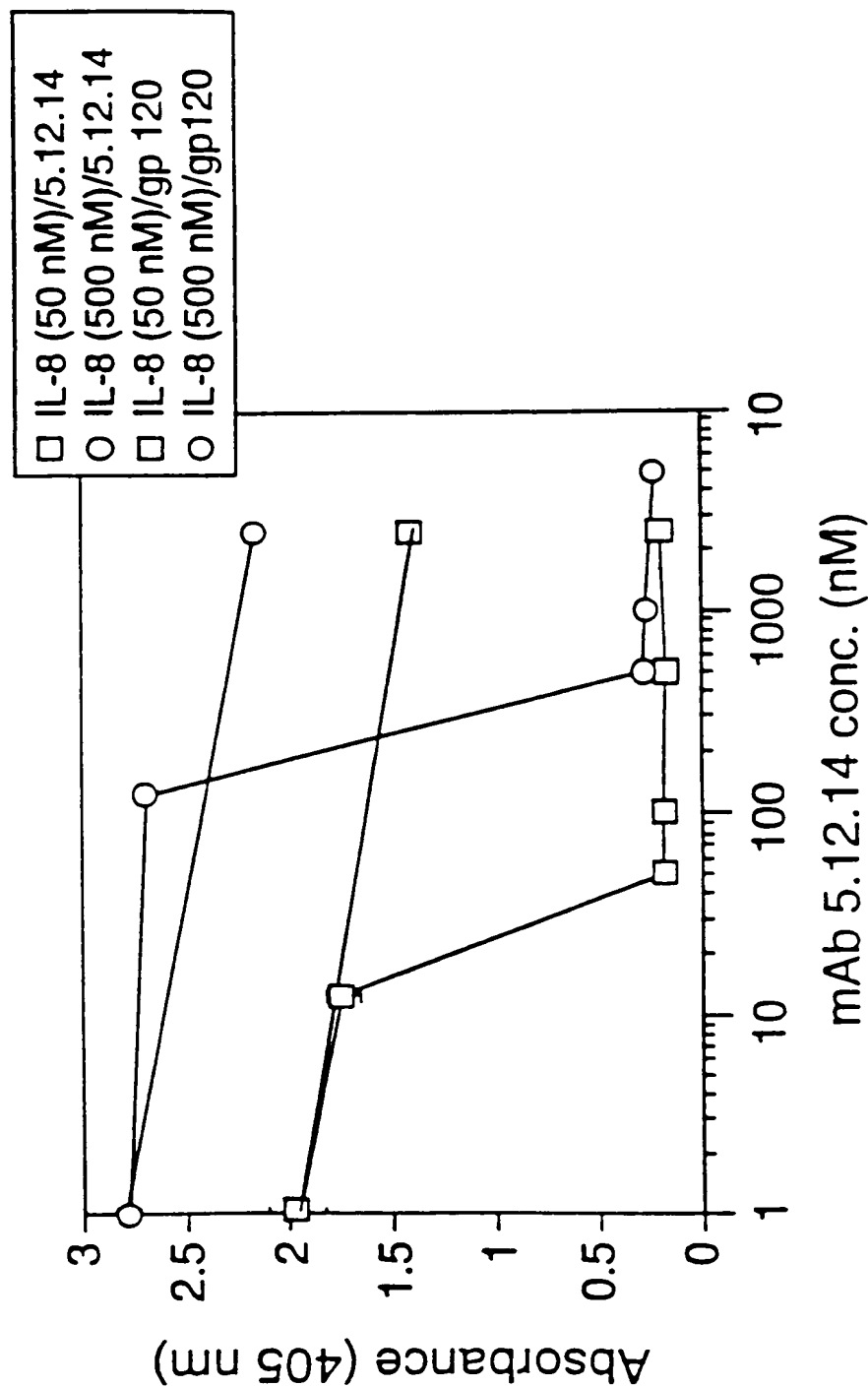




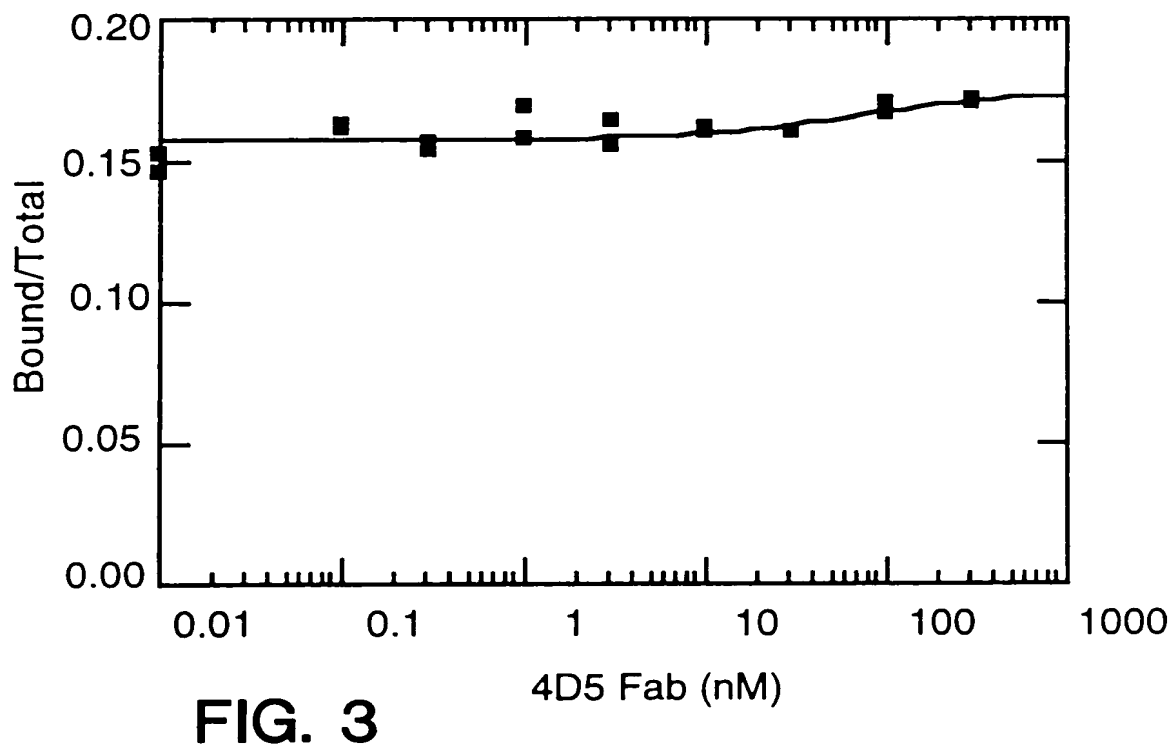
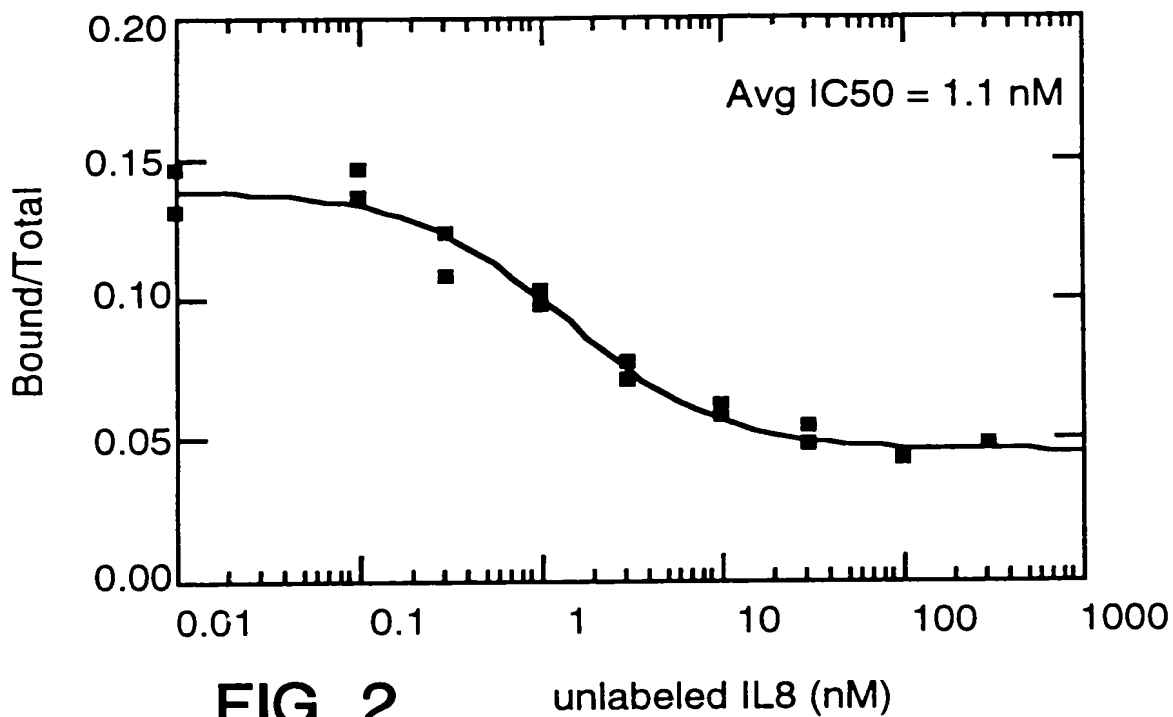
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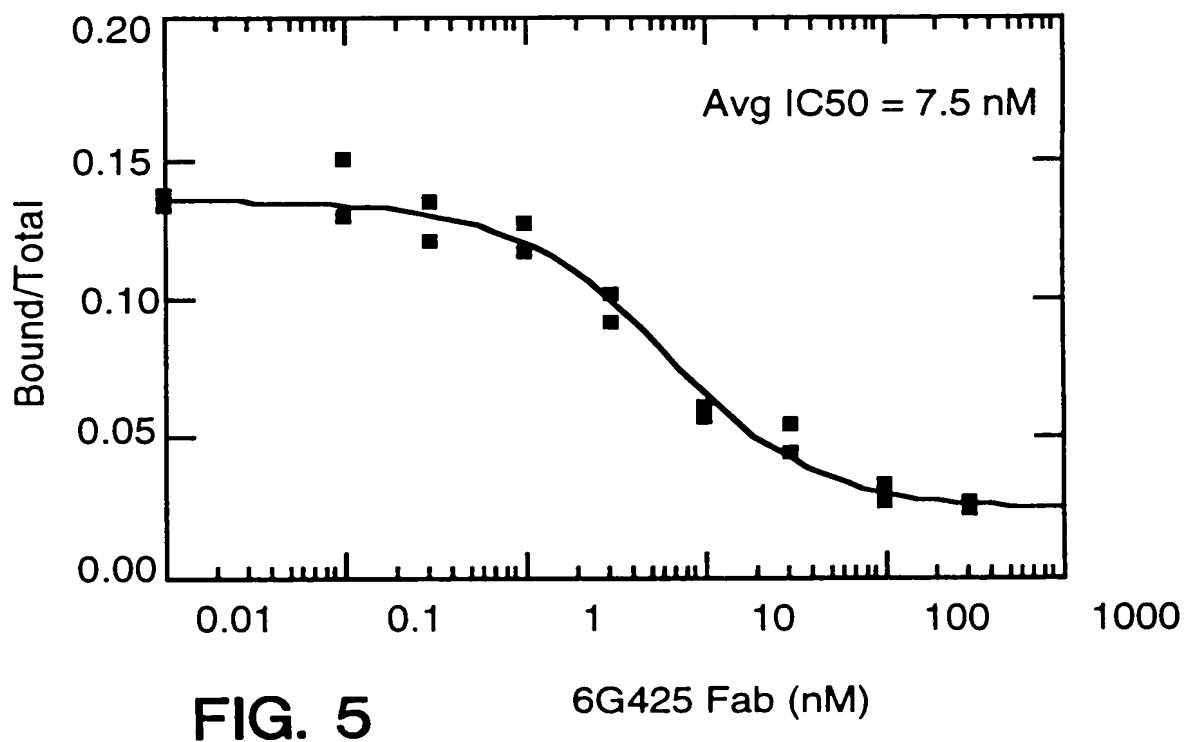
FIG. 1





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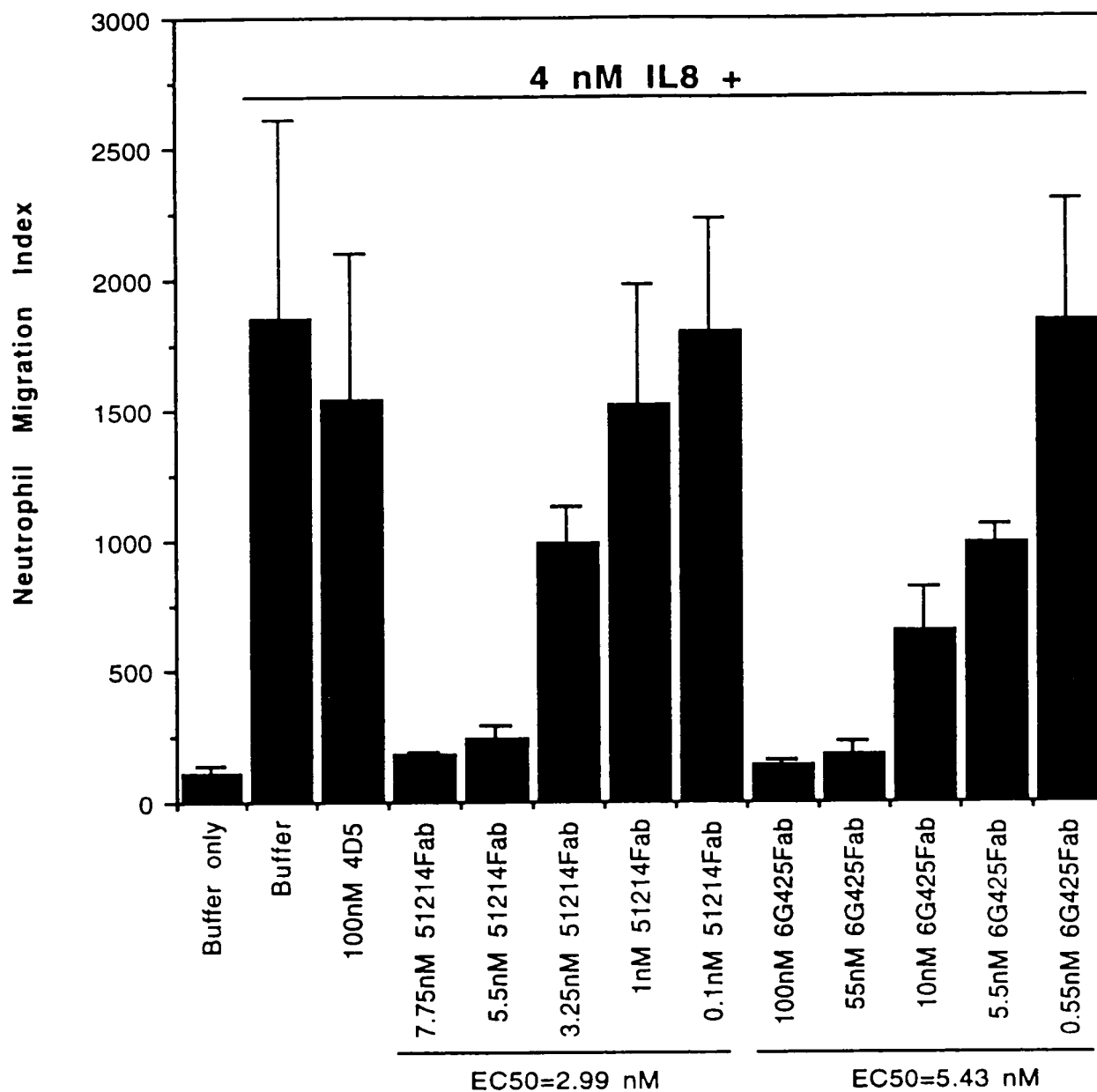


FIG. 6



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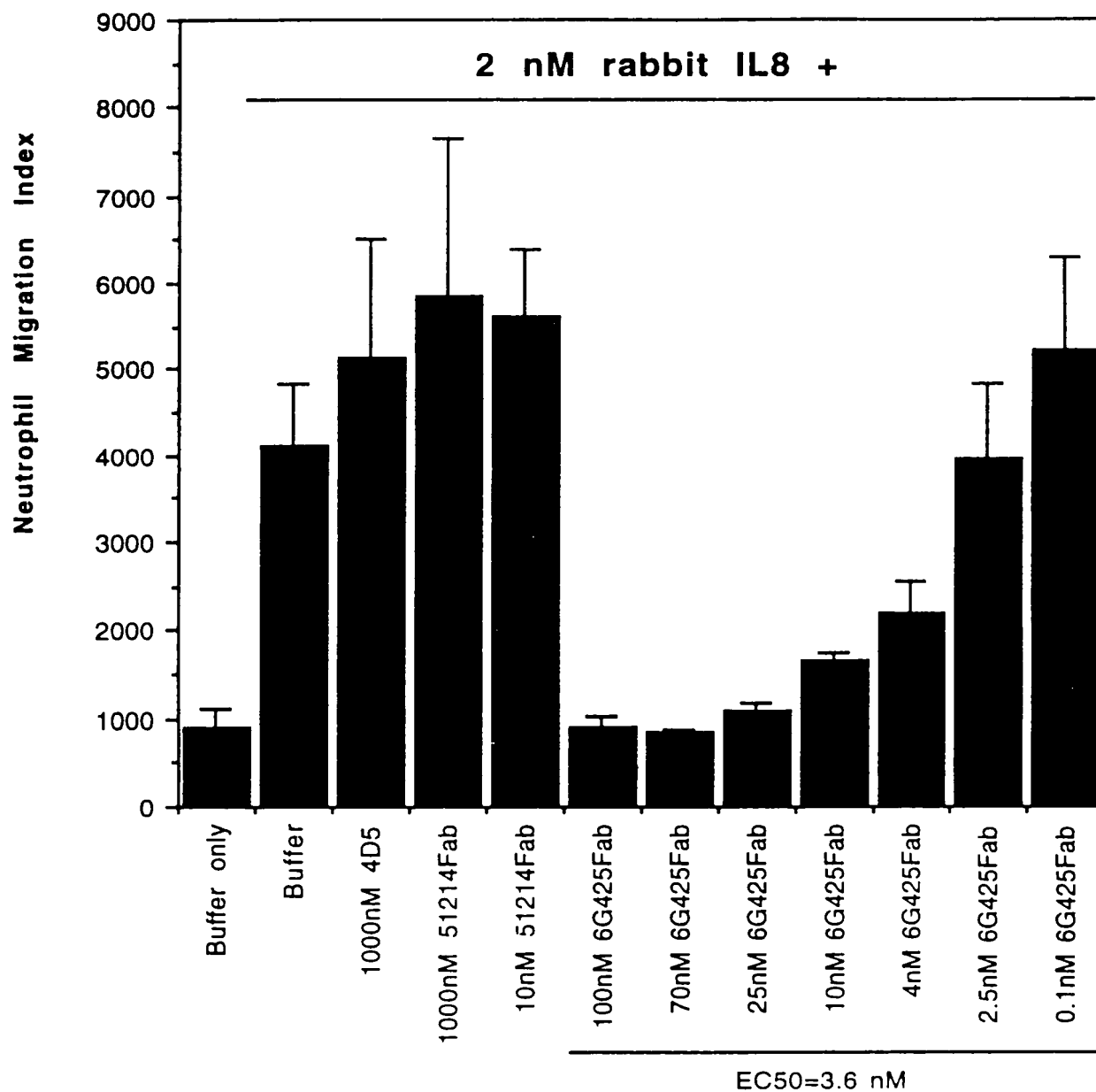
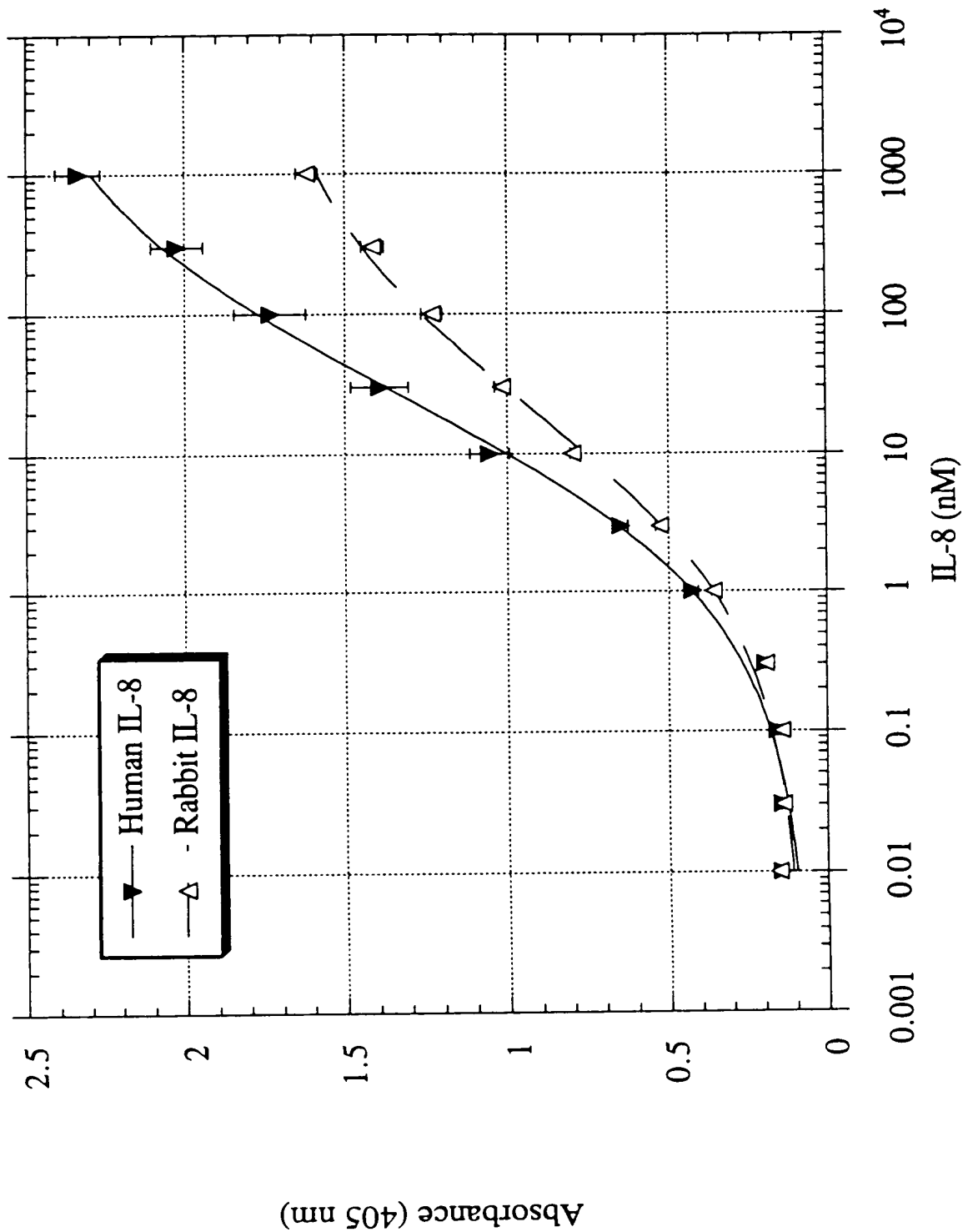


FIG. 7

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FIG. 8





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FIG. 9

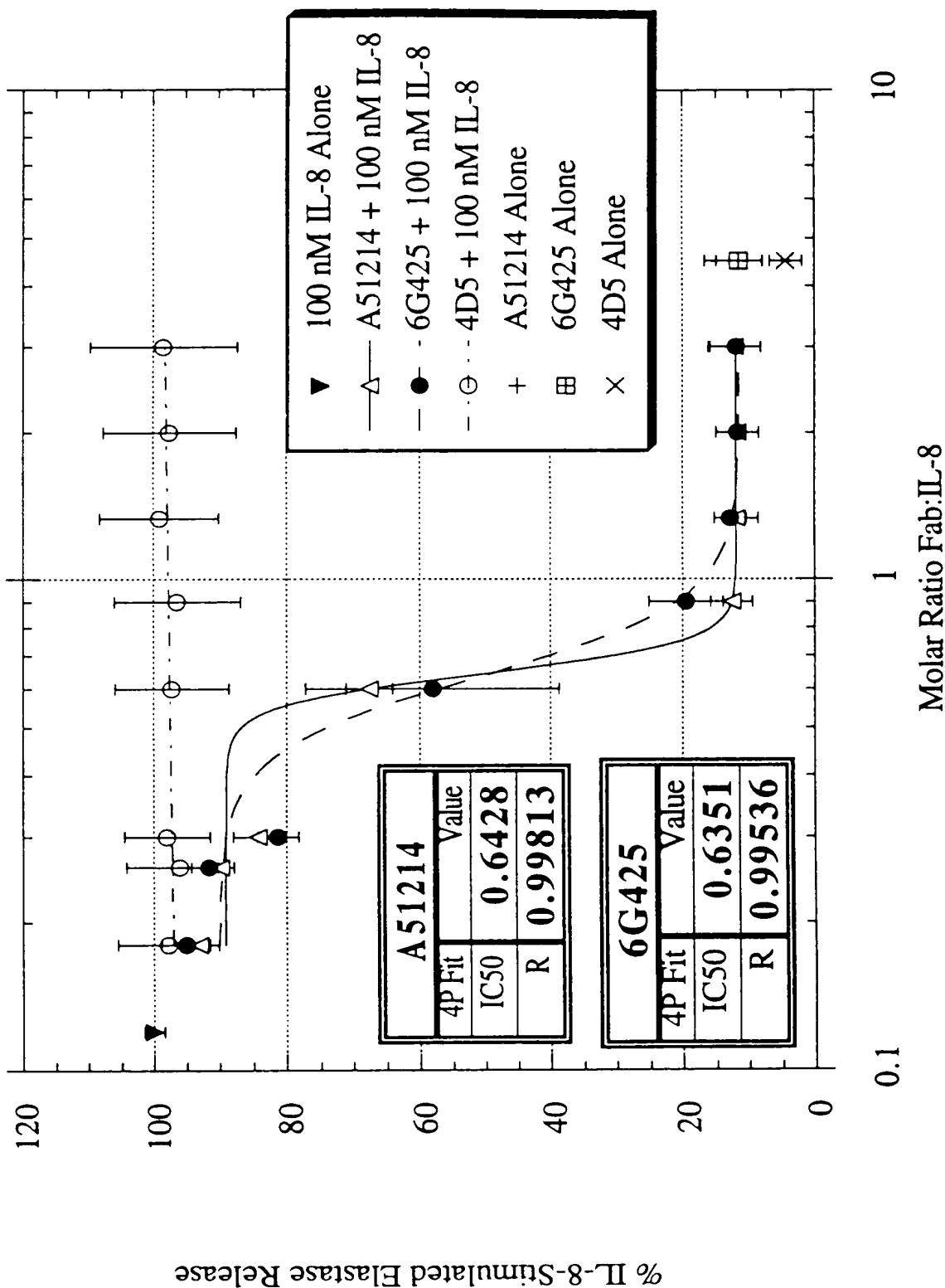
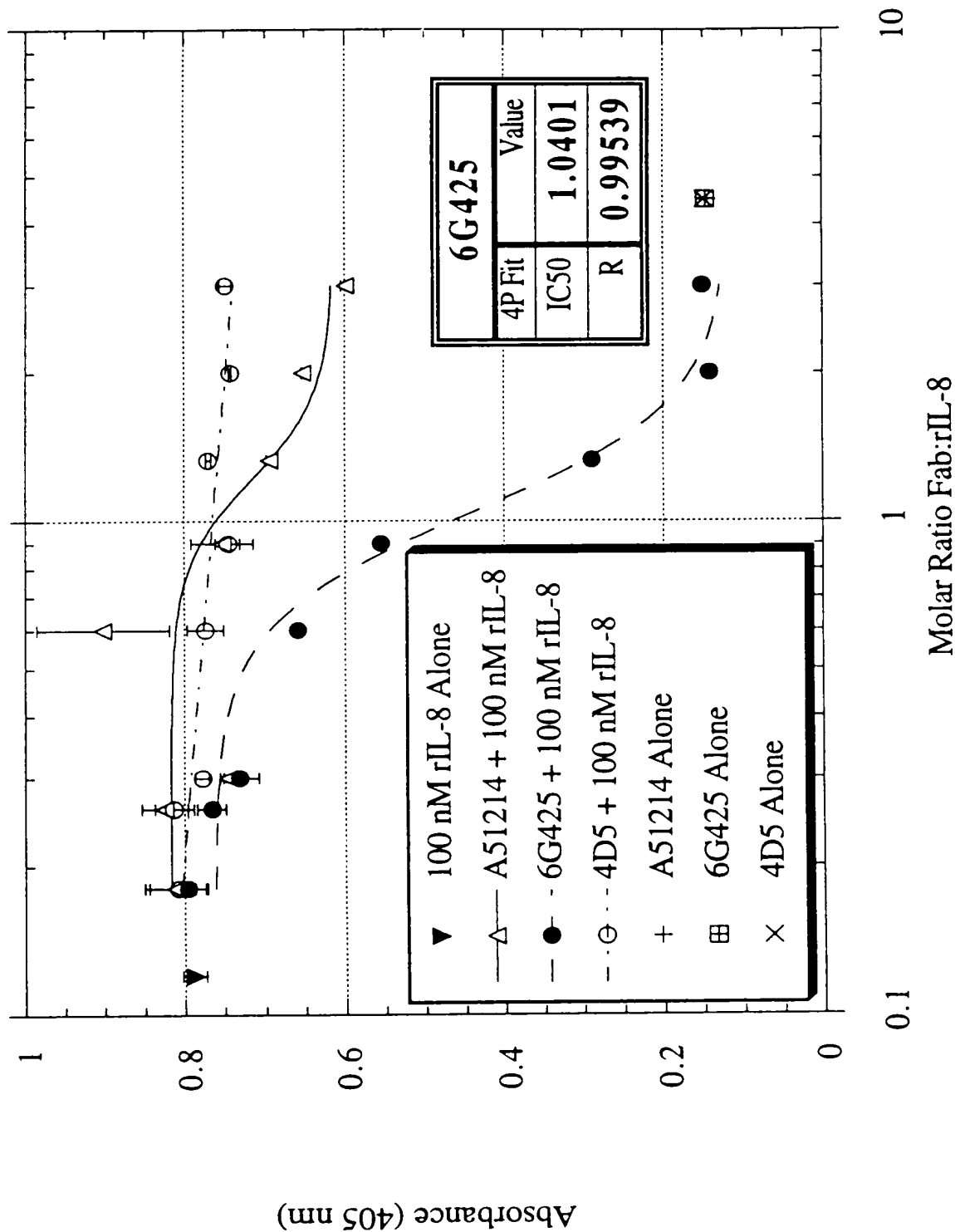




FIG. 10



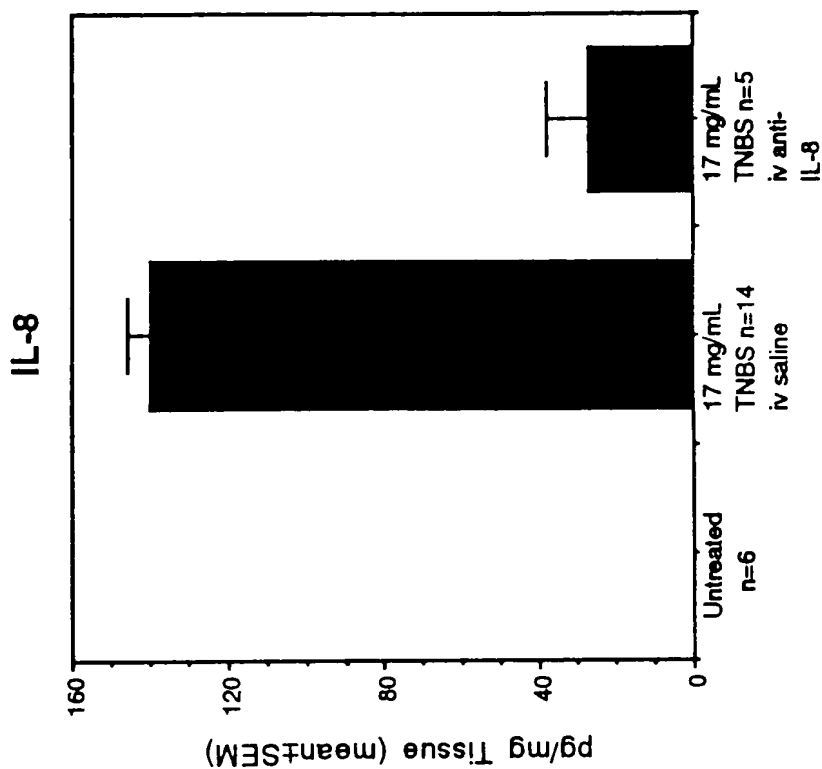


FIG. 11B

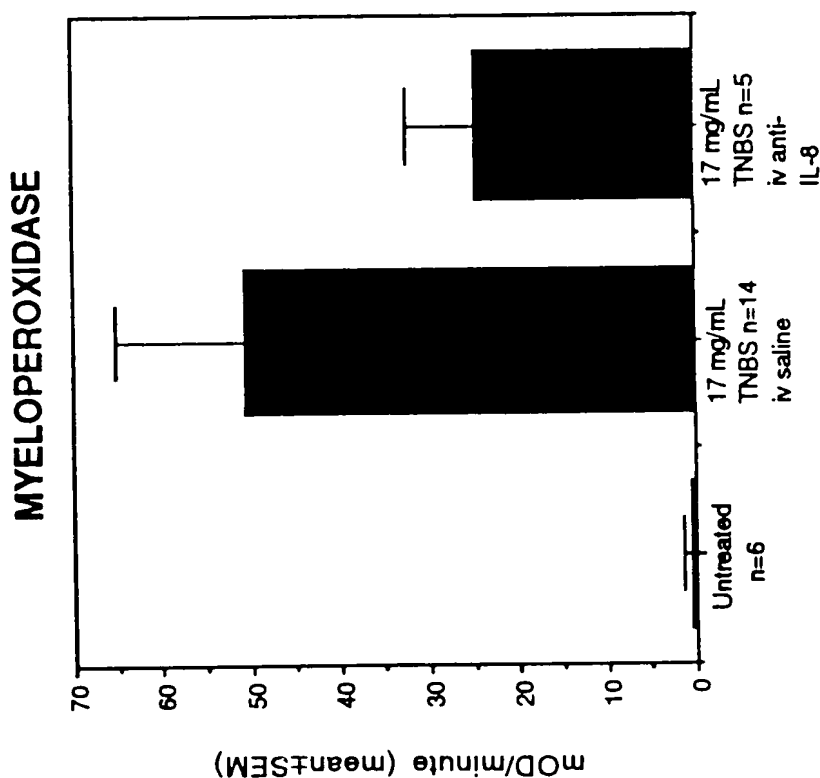


FIG. 11A

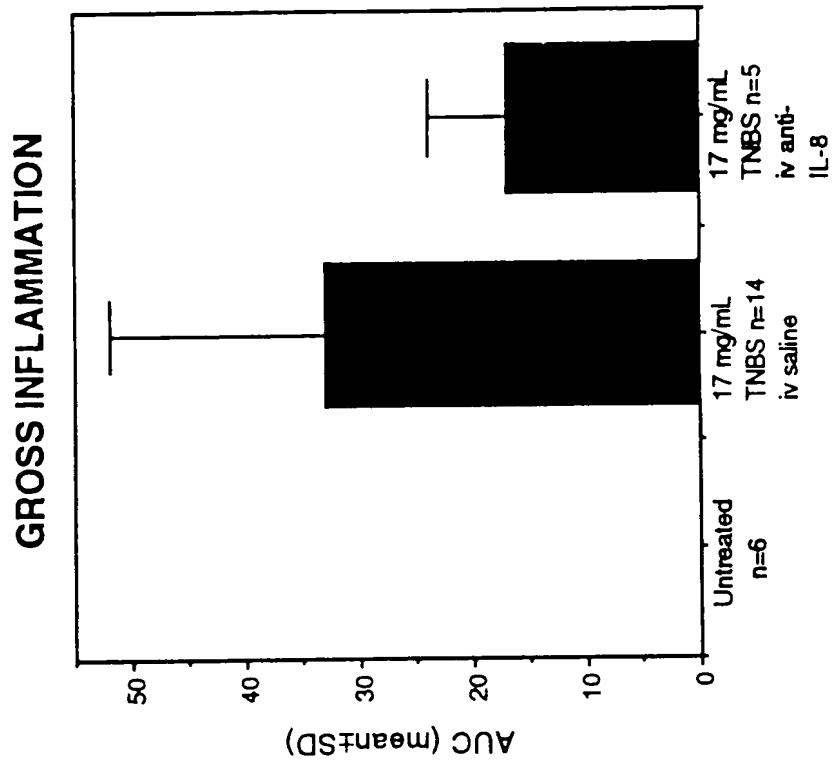
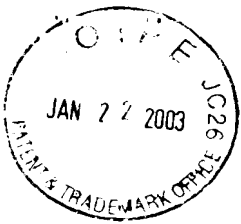


FIG. 11D

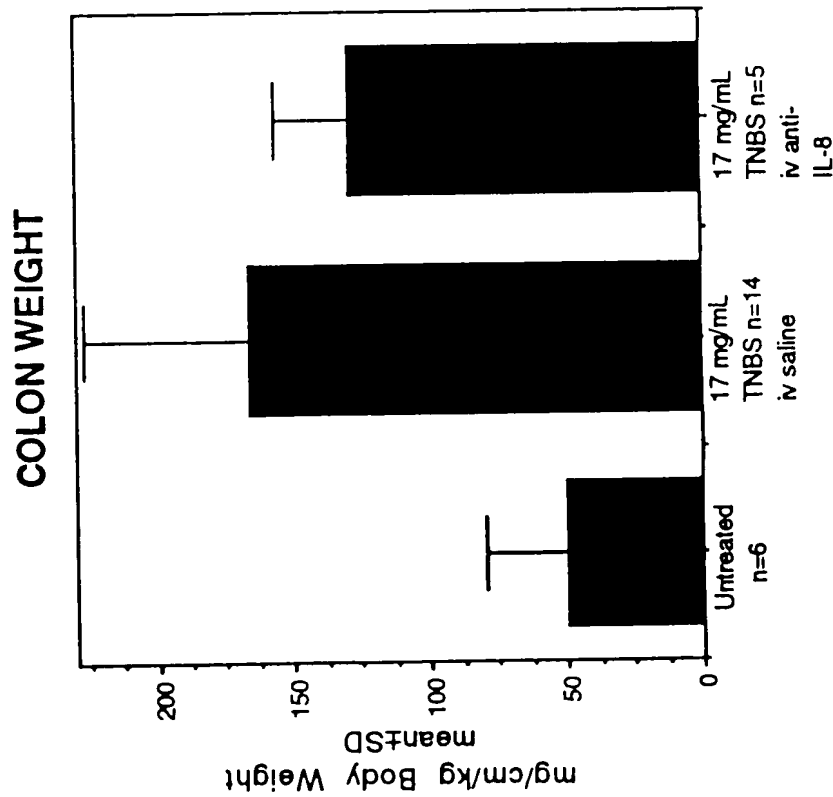


FIG. 11C

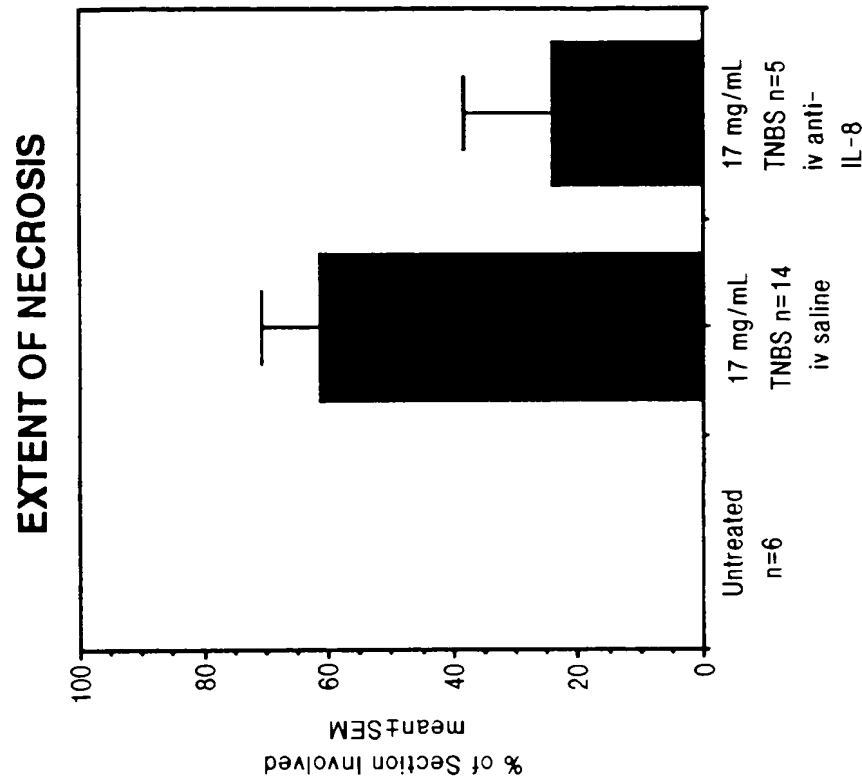


FIG. 11F

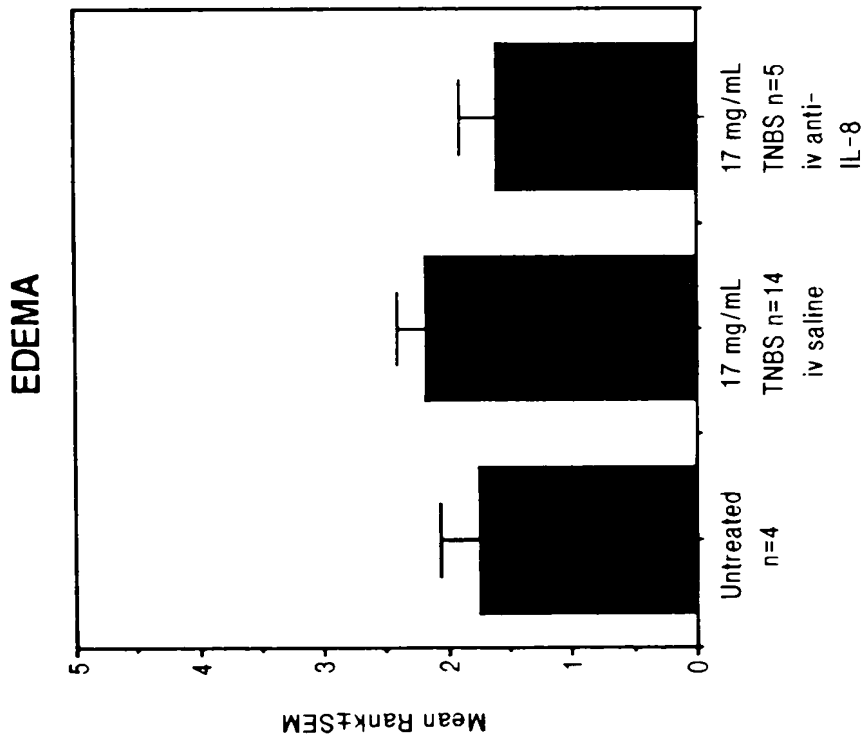


FIG. 11E

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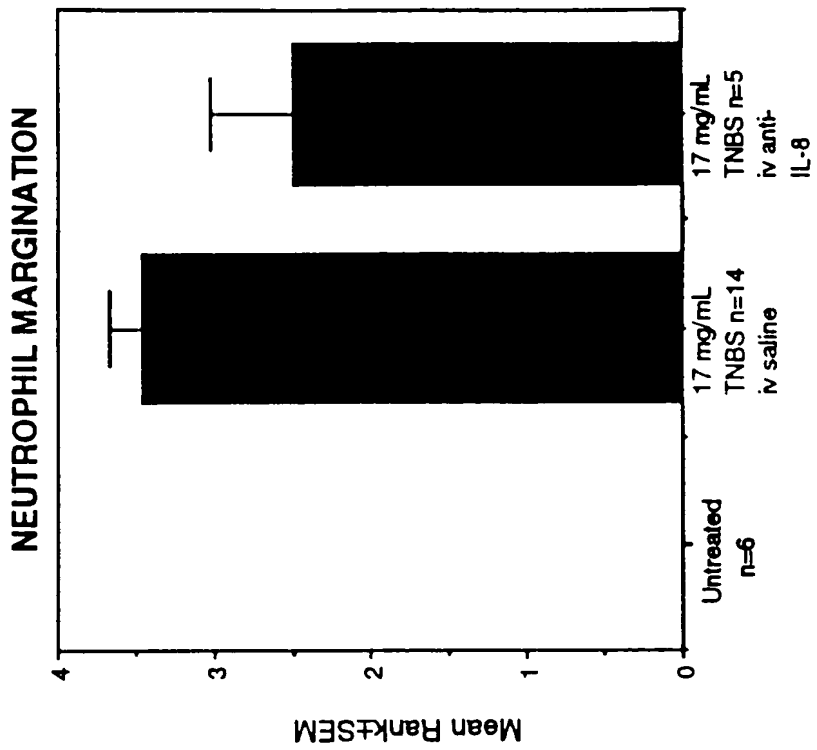


FIG. 11H

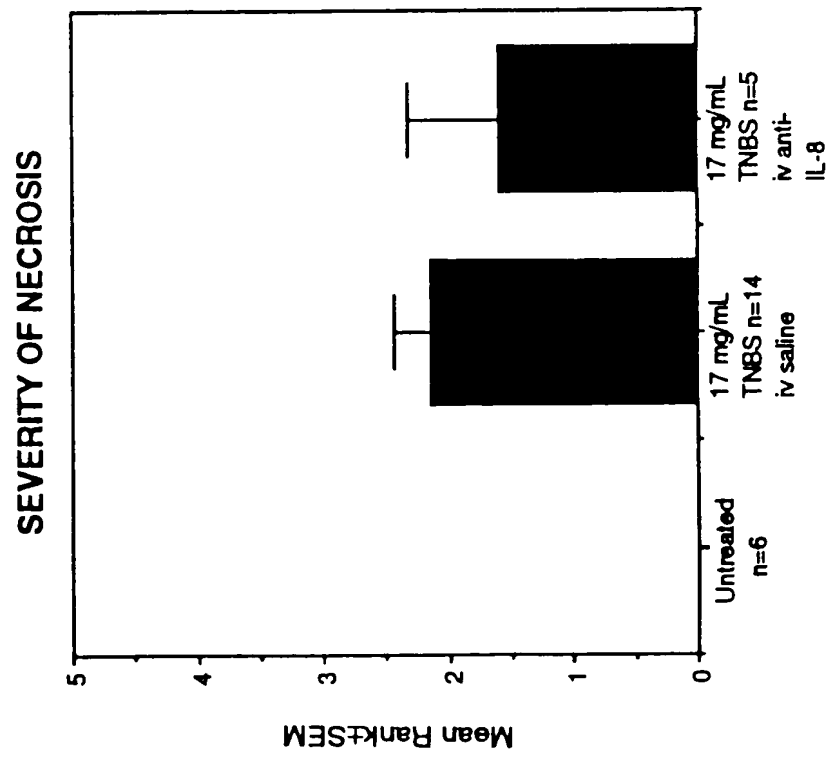


FIG. 11G



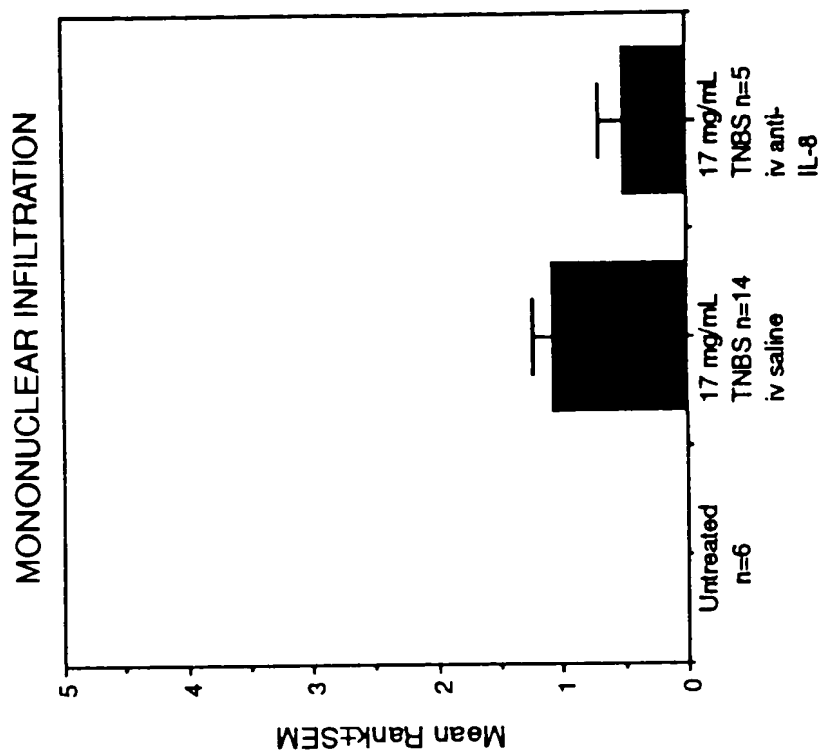


FIG. 11J

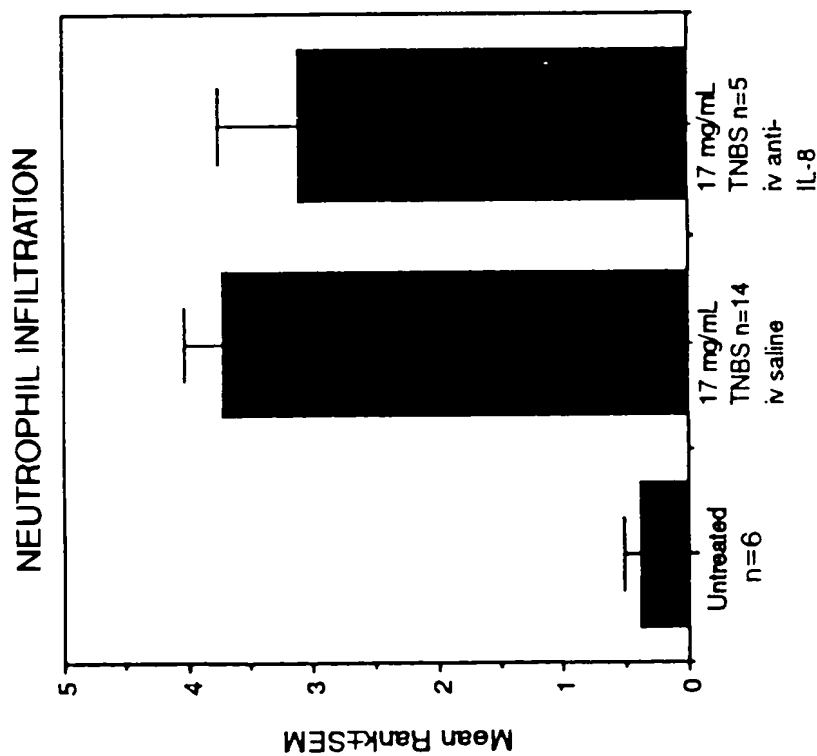


FIG. 11I





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FIG. 14

Light chain forward primer

SL001A-2 35 mer

5' ACAACGCGTAGCT GACATCGTCATGACCCAGTC 3' (SEQ ID NO.7)
T T T
A (SEQ ID NO.8)
(SEQ ID NO.9)

Light chain reverse primer

SL001B 37 mer

5' GCTCTTCGAATG GTGGGAAGATGGATACAGTTGGTGC 3' (SEQ ID NO.10)



FIG. 15

Heavy chain forward primer

SL002B 39 mer

5' CGATGGGCCCCG ATAGACCGATGGGGCTGTTGTTTGGC 3' (SEQ ID NO.11)
T (SEQ ID NO.12)
G (SEQ ID NO.13)
A (SEQ ID NO.14)

Heavy chain reverse primer

SL002B 39-MER

5' CGATGGGCCCCG ATAGACCGATGGGGCTGTTGTTTGGC 3' (SEQ ID NO.11)
T (SEQ ID NO.15)
A (SEQ ID NO.14)
G (SEQ ID NO.13)



1 GACATTGTCA TGACACAGTC TCAAAATTC ATGTCCACAT CAGTAGGAGA CAGGGTCAGC
CTGTAACAGT ACTGTGTCAG AGTTTTTAAG TACAGGTGTA GTCATCCTCT GTCCCAGTCG
1 D I V M T Q S Q K F M S T S V G D R V S

61 GTCACCTGCA AGCCAGTCA GAATGTGGGT ACTAATGTAG CCTGGTATCA ACAGAAACCA
CAGTGGACGT TCCGGTCAGT CTTACACCCA TGATTACATC GGACCATAGT TGTCTTTGGT
21 V T C K A S Q N V G T N V A W Y Q Q K P
* * * * * * * * * * * * *

CDR #1

121 GGGCAATCTC CTAAAGCACT GATTACTCG TCATCCTACC GGTACAGTGG AGTCCCTGAT
CCCGTTAGAG GATTTCGTGA CTAAATGAGC AGTAGGATGG CCATGTCACC TCAGGGACTA
41 G Q S P K A L I Y S S Y R Y S G V P D
* * * * * * * * * * * *

CDR #2

181 CGCTTCACAG GCAGTGGATC TGGACACAGT TTCACCTCTCA CCATCAGCCA TGTGCAGTCT
GGGAAGTGTC CGTCACCTAG ACCCTGTCTA AAGTGAGAGT GGTAGTCGGT ACACGTCAGA
61 R F T G S G S G T D F T L T I S H V Q S

241 GAAGACTTGG CAGACTATT CTGTCAGCAA TATAACATCT ATCCTCTCAC GTTCGGTCCT
CTTCTGAACC GTCTGATAA GACAGTCGTT ATATTGTAGA TAGGAGAGTG CAAGCCAGGA
81 E D L A D Y F C Q Q Y N I Y P L T F G P
* * * * * * * * * * * *

CDR #3

301 GGGACCAAGC TGGAGTTGAA ACGGGCTGAT GCTGCACCAC CAACTGTATC CATCTTCCCA
CCCTGGTTCG ACCTCAACTT TGCCCGACTA CGACGTGGTG GTTGACATAG GTAGAAGGCT
101 G T K L E L K R A D A A P P T V S I F P

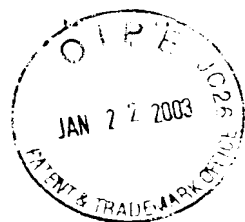
BstBI

361 CCATTTCGAA (SEQ ID NO.16)

GGTAAGCTT

121 P F E (SEQ ID NO.17)

FIG. 16



```
1  TTCTATTGCT  ACAAACGCGT  ACGCTGAGGT  GCAGCTGGTG  GAGTCTGGGG  GAGGCTTAGT
   AAGATAACGA  TGTTTGCGCA  TGCGACTCCA  CGTCGACCAC  CTCAGACCCC  CTCCGAATCA
1      E  V    Q  L  V    E  S  G  G    G  L  V

61  GCCGCCTGGA  GGGTCCCTGA  AACTCTCCTG  TGCAGCCTCT  GGATTTCATAT  TCAGTAGTTA
   CGGCGGACCT  CCCAGGGACT  TTGAGAGGAC  ACGTCGGAGA  CCTAAGTATA  AGTCATCAAT
13  P  P  G    G  S  L  K    L  S  C    A  A  S    G  F  I  F  S  S  Y
                                     *  *

                                   CDR #1

121  TGGCATGTCT  TGGGTTCGCC  AGACTCCAGG  CAAGAGCCTG  GAGTTGGTCG  CAACCATTAA
   ACCGTACAGA  ACCCAAGCGG  TCTGAGGTCC  GTTCTCGGAC  CTCAACCAGC  GTTGGTAATT
33  G  M  S    W  V  R  Q    T  P  G    K  S  L    E  L  V  A    T  I  N
   *  *  *                                     *  *  *

181  TAATAATGGT  GATAGCACCT  ATTATCCAGA  CAGTGTGAAG  GGCCGATTCA  CCATCTCCCG
   ATTATTACCA  CTATCGTGGA  TAATAGGTCT  GTCACACTTC  CCGGCTAAGT  GGTAGAGGGC
53  N  N  G  D  S  T  Y    Y  P  D    S  V  K    G  R  F  T    I  S  R
   *  *  *    *  *  *    *  *  *    *  *  *

                                   CDR #2

241  AGACAATGCC  AAGAACACCC  TGTACCTGCA  AATGAGCAGT  CTGAAGTCTG  AGGACACAGC
   TCTGTTACGG  TTCTTGTTGG  ACATGGACGT  TTA CTGTC A    GACTTCAGAC  TCCTGTGTCG
73  D  N  A    K  N  T  L    Y  L  Q    M  S  S    L  K  S  E    D  T  A

301  CATGTTTTAC  TGTGCAAGAG  CCCTCATTAG  TTCGGCTACT  TGGTTTGGTT  ACTGGGGCCA
   GTACAAAATG  ACACGTTCTC  GGGAGTAATC  AAGCCGATGA  ACCAAACCAA  TGACCCCGGT
93  M  F  Y    C  A  R  A    L  I  S  S  A  T  W  F  G  Y    W  G  Q
                                   *  *  *  *  *  *  *  *  *

                                   CDR #3

361  AGGGACTCTG  GTCACTGTCT  CTGCAGCCAA  AACAAACAGCC  CCATCTGTCT
   TCCCTGAGAC  CAGTGACAGA  GACGTCGGTT  TTGTTGTCGG  GGTAGACAGA
113  G  T  L    V  T  V  S    A  A  K    T  T  A    P  S  V  Y

      ApaI
411  ATCCGGG (SEQ ID NO.18)
      TAGGCC
130  P      (SEQ ID NO.19)
```

FIG. 17



Title: Antibody Fragment-Polymer Conjugates and Humanized Anti-IL-8

Monoclonal Antibodies; Inventor: Vanessa H. et al.

Application No.: 09 234,182 (Attorney Docket No. 09234.093A)

20 141

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FIG. 18

VL.front 31-MER

5' ACAACGCGTACGCTGATATCGTCATGACAG 3' (SEQ ID NO.20)

VL.rear 31-MER

5' GCAGCATCAGCTCTTCGAAGCTCCAGCTTGG 3' (SEQ ID NO.21)

VH.front.SPE 21-MER

5' CCACTAGTACGCAAGTTCACG 3' (SEQ ID NO.22)

VH.rear 33-MER

5' GATGGGCCCTTGGTGGAGGCTGCAGAGACAGTG 3' (SEQ ID NO.23)



1 ATGAAGAAGA ATATCGCATT TCTTCTTGCA TCTATGTTTCG TTTTCTCTAT TGCTACAAAC
TACTTCTTCT TATAGCGTAA AGAAGAACGT AGATACAAGC AAAAAAGATA ACGATGTTTG
-23 M K K N I A F L L A S M F V F S I A T N

61 GCGTACGCTG ATATCGTCAT GACACAGTCT CAAAAATTCA TGTCCACATC AGTAGGAGAC
CGCATGCGAC TATAGCAGTA CTGTGTCAGA GTTTTAAAGT ACAGGTGTAG TCATCCTCTG
-3 A Y A D I V M T Q S Q K F M S T S V G D

121 AGGGTCAGCG TCACCTGCAA GGCCAGTCAG AATGTGGGTA CTAATGTAGC CTGGTATCAA
TCCCAGTCGC AGTGGACGTT CCGGTCAGTC TTACACCCAT GATTACATCG GACCATAGTT
18 R V S V T C K A S Q N V G T N V A W Y Q
* * * * *
CDR #1

181 CAGAAACCAG GGCAATCTCC TAAAGCACTG ATTTACTCGT CATCCTACCG GTACAGTGGG
GTCTTTGGTC CCGTTAGAGG ATTTTCGTGAC TAAATGAGCA GTAGGATGGC CATGTCACCT
38 Q K P G Q S P K A L I Y S S S Y R Y S G
* * * * *
CDR #2

241 GTCCCTGATC GCTTCACAGG CAGTGGATCT GGGACAGATT TCACTCTCAC CATCAGCCAT
CAGGGACTAG CGAAGTGTCC GTCACCTAGA CCCTGTCTAA AGTGAGAGTG GTAGTCGGTA
58 V P D R F T G S G S G T D F T L T I S H

301 GTGCAGTCTG AAGACTTGGC AGACTATTTT TGTCAGCAAT ATAACATCTA TCCTCTCACG
CACGTCAGAC TTCTGAACCG TCTGATAAAG ACAGTCGTTA TATTGTAGAT AGGAGAGTGC
78 V Q S E D L A D Y F C Q Q Y N I Y P L T
* * * * *
CDR #3

BstBI

361 TTCGGTCCTG GGACCAAGCT GGAGCTTCGA AGAGCTGTGG CTGCACCATC TGTCTTCATC
AAGCCAGGAC CCTGGTTCGA CCTCGAAGCT TCTCGACACC GACGTGGTAG ACAGAAGTAG
98 F G P G T K L E L R R A V A A P S V F I

421 TTCCCGCCAT CTGATGAGCA GTTGAAATCT GGAAGTGTCT CTGTTGTGTG CCTGCTGAAT
AAGGGCGGTA GACTACTCGT CAACCTTTAGA CCTTGACGAA GACAACACAC GGACGACTTA
118 F P P S D E Q L K S G T A S V V C L L N

481 AACTTCTATC CCAGAGAGGC CAAAGTACAG TGGAAGGTGG ATAACGCCCT CCAATCGGGT
TTGAAGATAG GGTCTCTCCG GTTTCATGTC ACCTTCCACC TATTGCGGGA GGTTAGCCCA
138 N F Y P R E A K V Q W K V D N A L Q S G

541 AACTCCCAGG AGAGTGTACAC AGAGCAGGAC AGCAAGGACA GCACCTACAG CCTCAGCAGC
TTGAGGGTCC TCTCACAGTG TCTCGTCCTG TCGTTCCTGT CGTGGATGTC GGAGTCGTCTG
158 N S Q E S V T E Q D S K D S T Y S L S S

601 ACCCTGACGC TGAGCAAAGC AGACTACGAG AAACACAAAG TCTACGCCCTG CGAAGTCACC
TGGGACTGCG ACTCGTTTCG TCTGATGCTC TTTGTGTTTC AGATGCGGAC GCTTCAGTGG
178 T L T L S K A D Y E K H K V Y A C E V T

661 CATCAGGGCC TGAGCTCGCC CGTCACAAAG AGCTTCAACA GGGGAGAGTG
GTAGTCCCGG ACTCGAGCGG GCAGTGTTC TCGAAGTTGT CCCCTCTCAC
198 H Q G L S S P V T K S F N R G E C (SEQ ID NO.25)

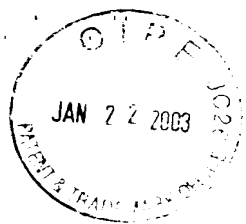
711 TTAA (SEQ ID NO.24)
AATT
216 O

FIG. 19



661 ACCCAGACCT ACATCTGCAA CGTGAATCAC AAGCCCAGCA ACACCAAGGT GGACAAGAAA
TGGGTCTGGA TGTAGACGTT GCACTTAGTG TTCGGGTCGT TGTGGTTCCA CCTGTTCTTT
198 T Q T Y I C N V N H K P S N T K V D K K
721 GTTGAGCCCA AATCTTGTGA CAAAACTCAC ACATGA (SEQ ID NO.26)
CAACTCGGGT TTAGAACACT GTTTTGAGTG TGTACT
218 V E P K S C D K T H T O (SEQ ID NO.27)

FIG. 20B



Title: Antibody Fragment-Polymer Conjugates and Humanized Anti-IL-8

Monoclonal Antibodies; Inventor: Vanessa H. et al.

Application No.: 09 234,182 (Attorney Docket No. 09234.093A)

24 141

Light Chain Primers:

MKLC-1, 22mer

5' CAGTCCAAGTGTTCAGGACGCC 3' (SEQ ID NO.1)

MKLC-2, 22mer

5' GTGCTGCTCATGCTGTAGGTGC 3' (SEQ ID NO.2)

MKLC-3, 23mer

5' GAAGTTGATGTCTTGTGAGTGGC 3' (SEQ ID NO.3)

Heavy Chain Primers:

IGG2AC-1, 24mer

5' GCATCCTAGAGTCACCGAGGAGCC 3' (SEQ ID NO.4)

IGG2AC-2, 22mer

5' CACTGGCTCAGGGAAATAACCC 3' (SEQ ID NO.5)

IGG2AC-3, 22mer

5' GGAGAGCTGGGAAGGTGTGCAC 3' (SEQ ID NO.6)

FIG. 21

Light chain forward primer

6G4.light.Nsi 36-MER

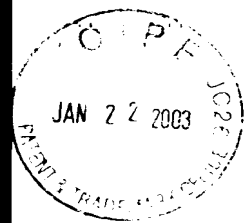
5'	CCAATGCATACGCT	GAC	ATC	GTG	ATG	ACC	CAG	ACC	CC	3'	(SEQ ID NO.28)
		T	T			T			T		(SEQ ID NO.29)
						A			A		(SEQ ID NO.30)

Light chain reverse primer

6G4.light.Mun 35-MER

5' AGA TGT CAA TTG CTC ACT GGA TGG TGG GAA GAT GG 3' (SEQ ID NO.31)

FIG. 22



Heavy chain forward primer

6G4.heavy.Mlu 32-MER

5' CAAACGCGTAGCT GAG ATC CAG CTG CAG CAG 3' (SEQ ID NO.32)
T C (SEQ ID NO.33)

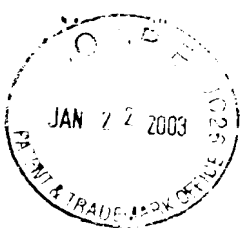
Heavy chain reverse primer

SL002B 39-MER

5' CGATGGGCCCCG ATAGACCGATGGGGCTGTTGTTTGGC 3' (SEQ ID NO.11)
T (SEQ ID NO.15)
A (SEQ ID NO.14)
G (SEQ ID NO.13)

FIG. 23





5' CTTGGTGGAGGCGGAGGAGACG 3' (SEQ ID NO.38)

Mutagenesis Primer for 6G425VL

DS/VF 38MER

5' GAAACGGGCTGTTGCTGCACCAACTGTATTCATCTTCC 3' (SEQ ID NO.39)

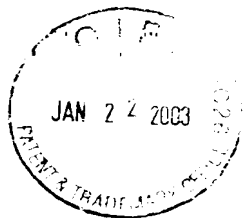
SYN.BstEII 31 MER

5' GTCACCGTCT CCTCCGCCTC CACCAAGGGC C 3' (SEQ ID NO.40)

SYN.Apa 22 MER

5' CTTGGTGGAGGCGGAGGAGACG 3' (SEQ ID NO.38)

FIG. 26



Title: Antibody Fragment-Polymer Conjugates and Humanized Anti-IL-8

Monoclonal Antibodies; Inventor: Vanessa H. et al.

Application No.: 09 234,182 (Attorney Docket No. C-9219-093A)

31 141

661 GCCTGCGAAG TCACCCATCA GGGCCTGAGC TCGCCCGTCA CAAAGAGCTT CAACAGGGGA
CGGACGCTTC AGTGGGTAGT CCGGACTCG AGCGGGCAGT GTTCTCGAA GTTGTCCTT
198 A C E V T H Q G L S S P V T K S F N R G

721 GAGTGTAA (SEQ ID NO.41)

CTCACAATT

218 E C O (SEQ ID NO.42)

FIG. 27B


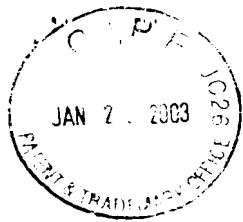
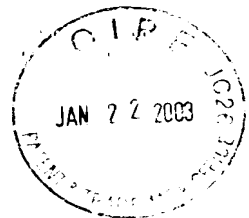


FIG. 28A



661 TTGGGCACCC AGACCTACAT CTGCAACGTG AATCACAAGC CCAGCAACAC CAAGGTGGAC
AACCCGTGGG TCTGGATGTA GACGTTGCAC TTAGTGTTCG GGTGTTGTG GTTCCACCTG
198 L G T Q T Y I C N V N H K P S N T K V D
721 AAGAAAGTTG AGCCCAAATC TTGTGACAAA ACTCACACAT GA (SEQ ID NO.43)
TTCTTTCAAC TCGGGTTTAG AACACTGTTT TGAGTGTGTA CT
218 K K V E P K S C D K T H T O (SEQ ID NO.44)

FIG. 28B



Variable Light Chain Domain

```

      10      20      abcde 30      40
6G425  DIVMTQTPLSLPVS LGDQASISCRSSQSLVHGIGNTYLHWYLQKPGQSPKLLIY
      * * * * *
F(ab)-1 DIQMTQSPSSLSASVGDRTITCRSSQSLVHGIGNTYLHWYQQKPGKAPKLLIY
      * * * * *
humkI   DIQMTQSPSSLSASVGDRTITCRASKTI-----SKYLAWYQQKPGKAPKLLIY
      =====
      ++++++
      L1

      50      60      70      80      90      100
6G425  YKVSNRFGVFPDRFSDSGSGTDFTLRISRVEAEDLGLYFCSQSTHVPLTFGAGTKLELKR (SEQ ID NO.45)
      * * * * *
F(ab)-1 YKVSNRFGVPSRFSGSGSGTDFTLTISLQPEDFATYYCSQSTHVPLTFGQGTKVEIKR (SEQ ID NO.46)
      * * * * *
humkI   YSGSTLESQVPSRFSGSGSGTDFTLTISLQPEDFATYYCQQHNEYPLTFGQGTKVEIKR (SEQ ID NO.47)
      ===
      ++++++
      L2
      ++++++
      L3
```

Variable Heavy Chain Domain

```

      10      20      30      40
6G425  EIQLQQSGPELMKPGASVKISCKASGYSFSSHYMHVWKQSHGKSLEWI
      * * * * *
F(ab)-1 EVQLVESGGGLVQPGGSLRLSCAASGYSFSSHYMHVVRQAPGKGLEWV
      * * * * *
humIII  EVQLVESGGGLVQPGGSLRLSCAASGFSFTGHWMNWRQAPGKGLEWV
      =====
      +++++
      H1

      50  a      70      80  abc      90      100      110
6G425  GYIDPSNGETTYNQKFKGKATLTVDTSSTANVHLSSLTSDDSAVYFCAARGDYRYNGDWFFDVWGAGT (SEQ ID NO.48)
      * * * * *
F(ab)-1 GYIDPSNGETTYNQKFKGRFTISRDNKNTLYLQMNSLRAEDTAVYYCAARGDYRYNGDWFFDVWGQGT (SEQ ID NO.49)
      * * * * *
humIII  GMIHPSDSETRYADSVKGRFTISRDNKNTLYLQMNSLRAEDTAVYYCAARGIYFY-GTTYFDYWQGT (SEQ ID NO.50)
      ===
      ++++++
      H2
      ++++++
      H3
```

FIG. 29

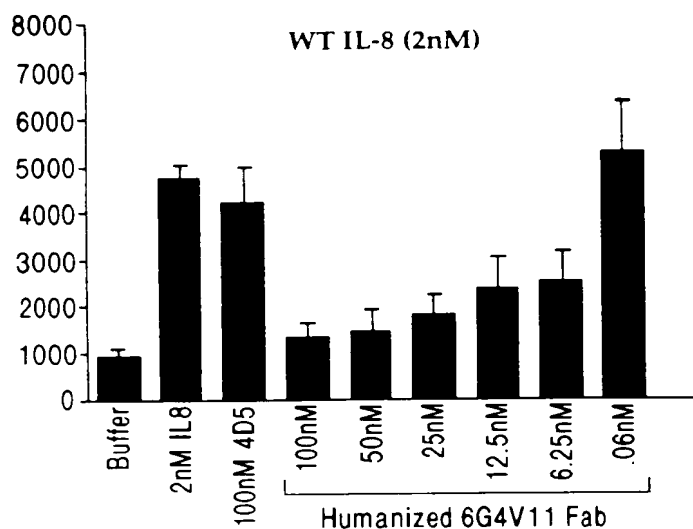


FIG. 30A

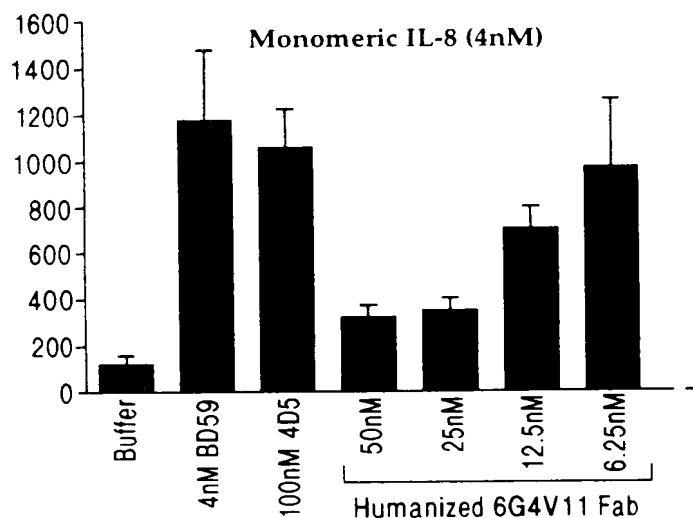


FIG. 30B

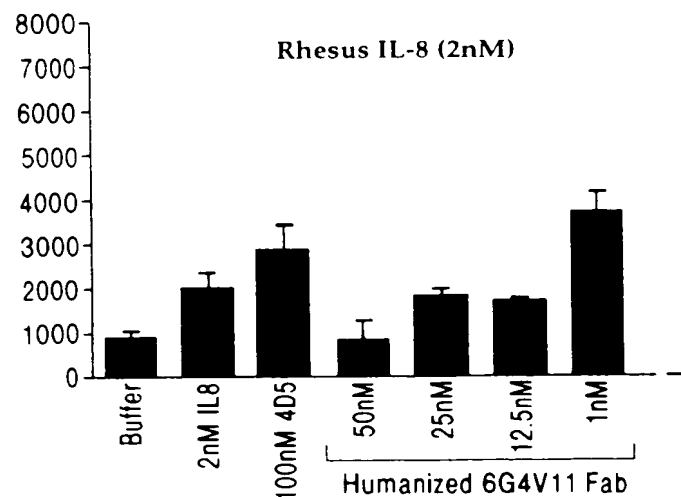
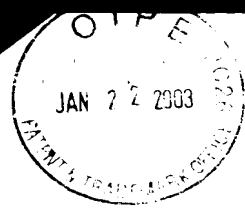


FIG. 30C



1 ATGAAAAAGA ATATCGCATT TCTTCTTGCA TCTATGTTCG TTTTCTCTAT TGCTACAAAC
TACTTTTTCT TATAGCGTAA AGAAGAACGT AGATACAAGC AAAAAAGATA ACGATGTTTG
-23 M K K N I A F L L A S M F V F S I A T N

61 GCATACGCTG ATATCCAGAT GACCCAGTCC CCGAGCTCCC TGTCCGCCTC TGTGGGCGAT
CGTATGCGAC TATAGGTCTA CTGGGTCAGG GGCTCGAGGG ACAGGCGGAG ACACCCGCTA
-3 A Y A D I Q M T Q S P S S L S A S V G D

121 AGGGTCACCA TCACCTGCAG GTCAAGTCAA AGCTTAGTAC ATGGTATAGG TAACACGTAT
TCCCAGTGGT AGTGGACGTC CAGTTCAGTT TCGAATCATG TACCATATCC ACGATGCATA
18 R V T I T C R S S Q S L V H G I G N T Y

181 TTACACTGGT ATCAACAGAA ACCAGGAAAA GCTCCGAAAC TACTGATTTA CAAAGTATCC
AATGTGACCA TAGTTGTCTT TGGTCCTTTT CGAGGCTTTG ATGACTAAAT GTTTCATAGG
38 L H W Y Q Q K P G K A P K L L I Y K V S

241 AATCGATTCT CTGGAGTCCC TTCTCGCTTC TCTGGATCCG GTTCTGGGAC GGATTTCACT
TTAGCTAAGA GACCTCAGGG AAGAGCGAAG AGACCTAGGC CAAGACCCTG CCTAAAGTGA
58 N R F S G V P S R F S G S G S G T D F T

301 CTGACCATCA GCAGTCTGCA GCCAGAAGAC TTCGCAACTT ATTACTGTTC ACAGAGTACT
GACTGGTAGT CGTCAGACGT CGGTCTTCTG AAGCGTTGAA TAATGACAAG TGTCTCATGA
78 L T I S S L Q P E D F A T Y Y C S Q S T

361 CATGTCCCGC TCACGTTTGG ACAGGGTACC AAGGTGGAGA TCAAACGAAC TGTGGCTGCA
GTACAGGGCG AGTGCAAACC TGTCCTCATG TTCCACCTCT AGTTTGCTTG ACACCGACGT
98 H V P L T F G Q G T K V E I K R T V A A

421 CCATCTGTCT TCATCTTCCC GCCATCTGAT GAGCAGTTGA AATCTGGAAC TGCTTCTGTT
GGTAGACAGA AGTAGAAGGG CGGTAGACTA CTCGTCAACT TTAGACCTTG ACGAAGACAA
118 P S V F I F P P S D E Q L K S G T A S V

481 GTGTGCCTGC TGAATAACTT CTATCCCAGA GAGGCCAAAG TACAGTGGAA GGTGGATAAC
CACACGGACG ACTTATTGAA GATAGGGTCT CTCCGGTTTC ATGTCACCTT CCACCTATTG
138 V C L L N N F Y P R E A K V Q W K V D N

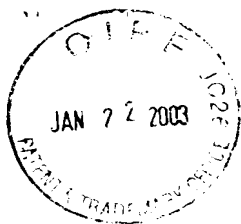
541 GCCCTCCAAT CGGGTAACTC CCAGGAGAGT GTCACAGAGC AGGACAGCAA GGACAGCACC
CGGGAGGTTA GCCATTGAG GGTCCTCTCA CAGTGTCTCG TCCTGTCTGTT CCTGTCTGTTG
158 A L Q S G N S Q E S V T E Q D S K D S T

601 TACAGCCTCA GCAGCACCTT GACGCTGAGC AAAGCAGACT ACGAGAAACA CAAAGTCTAC
ATGTCGGAGT CGTCGTGGGA CTGCGACTCG TTTCGTCTGA TGCTCTTTGT GTTTCAGATG
178 Y S L S S T L T L S K A D Y E K H K V Y

661 GCCTGCGAAG TCACCCATCA GGGCCTGAGC TCGCCCGTCA CAAAGAGCTT CAACAGGGGA
CGGACGCTTC AGTGGGTAGT CCCGGACTCG AGCGGGCAGT GTTTCTCGAA GTTGTCCCCT
198 A C E V T H Q G L S S P V T K S F N R G

721 GAGTGTTAAG CTGATCCTCT ACGCCGGACG CATCGTGGCC CTAGTACGCA ACTAGTCGTA
CTCACAATTC GACTAGGAGA TGCGGCCTGC GTAGCACCGG GATCATGCGT TGATCAGCAT
218 E C O (SEQ ID NO.51)

FIG. 31B



Amino Acid Sequence of the humanized anti-IL-8 6G4.2.5V19 Light Chain

MKKNIAFLASMFVFSIATNAYADIQMTQSPSSLSASVGDRTTITCRSSQSLVHGIGNTY
LHWYQQKPGKAPKLLIYKVSINRFSGVPSRFSGSGGTDFLTITSSLPEDFATYYCSQST
HVP LTFGGQGTKVEIKRTVAAPSVFIFPPSDEQLKSGTASVCLLNNFYPREAKVQWKVDN
ALQSGNSQESVTEQDSKDSYSTLSSTLTLSKADYEKHKVYACEVTHQGLSSPVTKSFNRG
EC (SEQ ID NO.51)

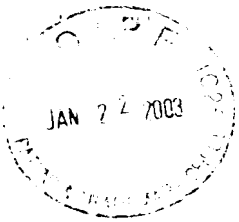
Amino Acid Sequence of the humanized anti-IL-8 6G4.2.5V19 Heavy Chain

MKKNIAFLASMFVFSIATNAYAEVQLVESGGGLVQPGGSLRLSCAASGYSFSSHYMH
WVKQAPGKGLEWVGVIDPSNGETTYNQKFKGRFTLSRDNSKNTAYLQMNSLRAEDTAVYY
CARGDYRYNGDWFFDVGQGT LVTVSSASTKGPSVFPLAPSSKSTSGGTAALGCLVKDYF
PEPVTWNSGALTSGVHTFPAVLQSSGLYSLSSVTVTPSSSLGTQTYICNVNHHKPSNTK
VDKKVEPKSCDKTHT (SEQ ID NO.55)

FIG. 31C



FIG. 32



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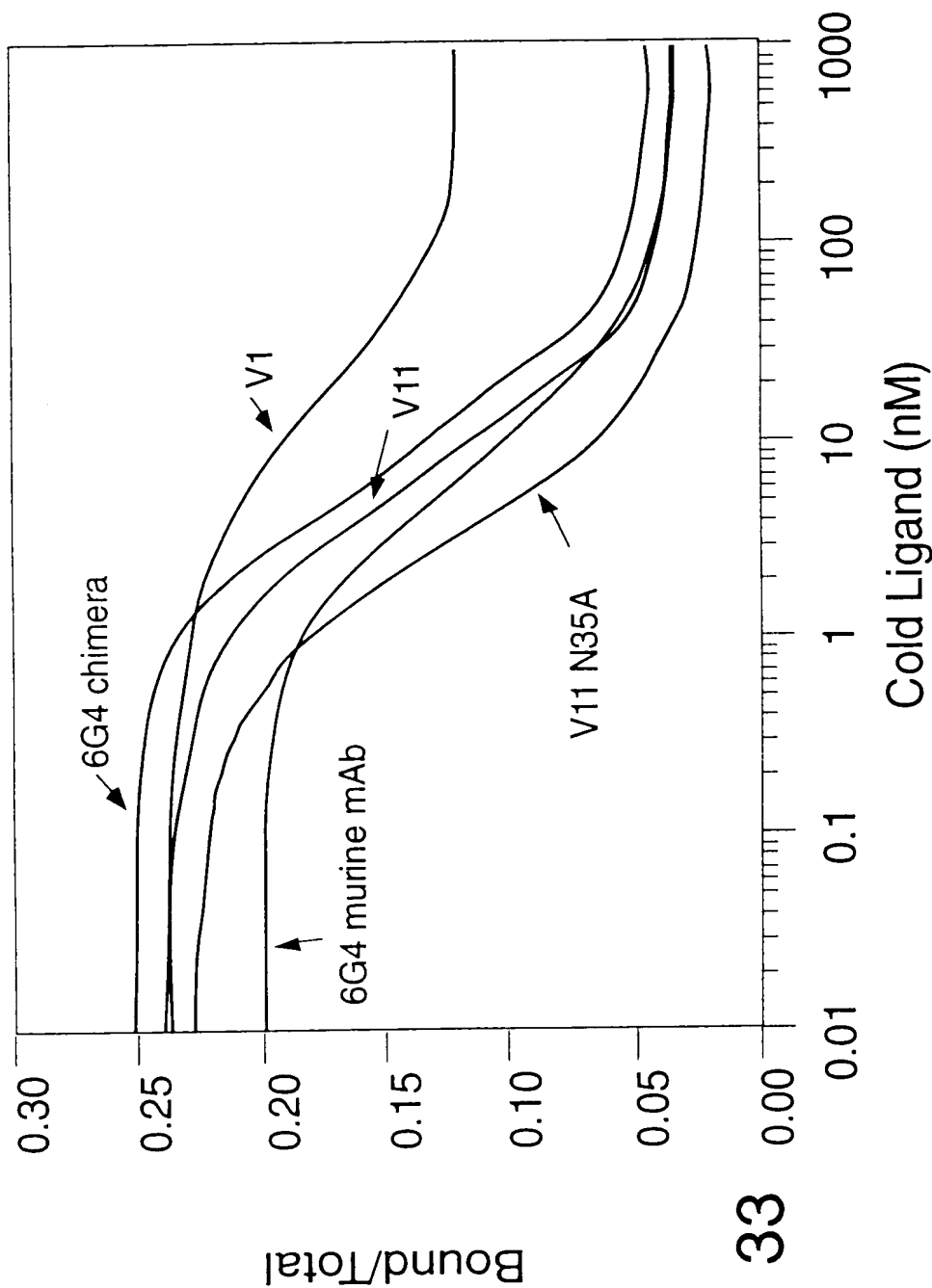
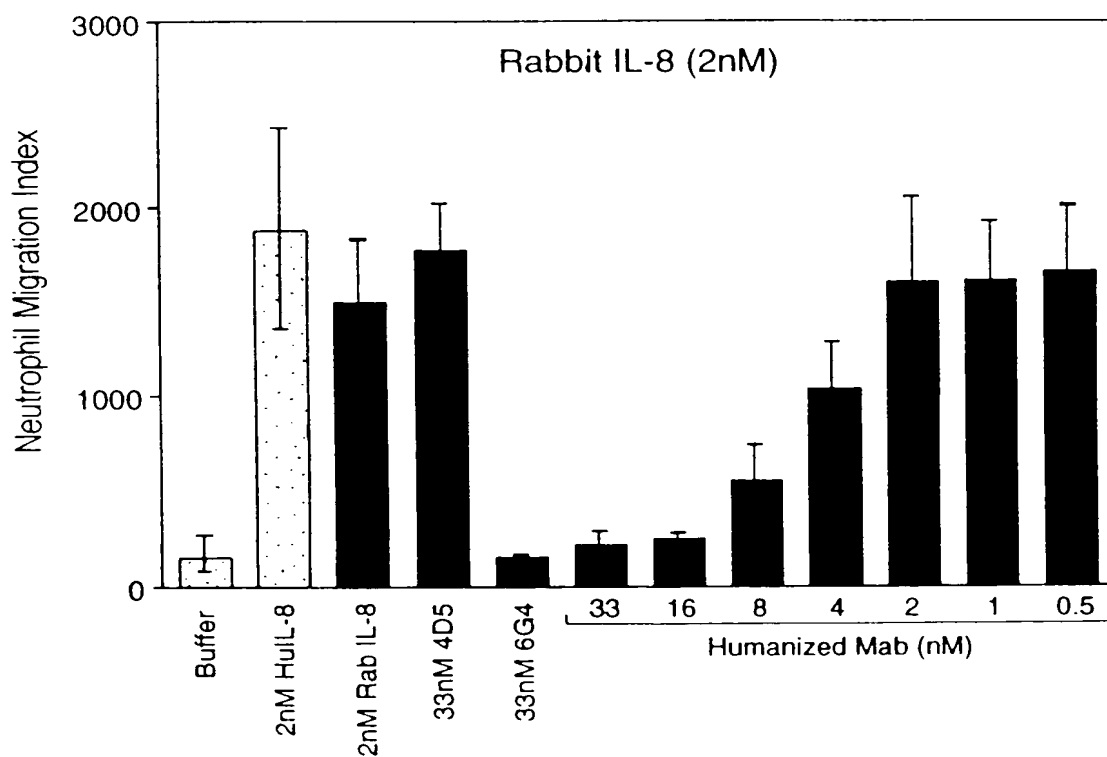
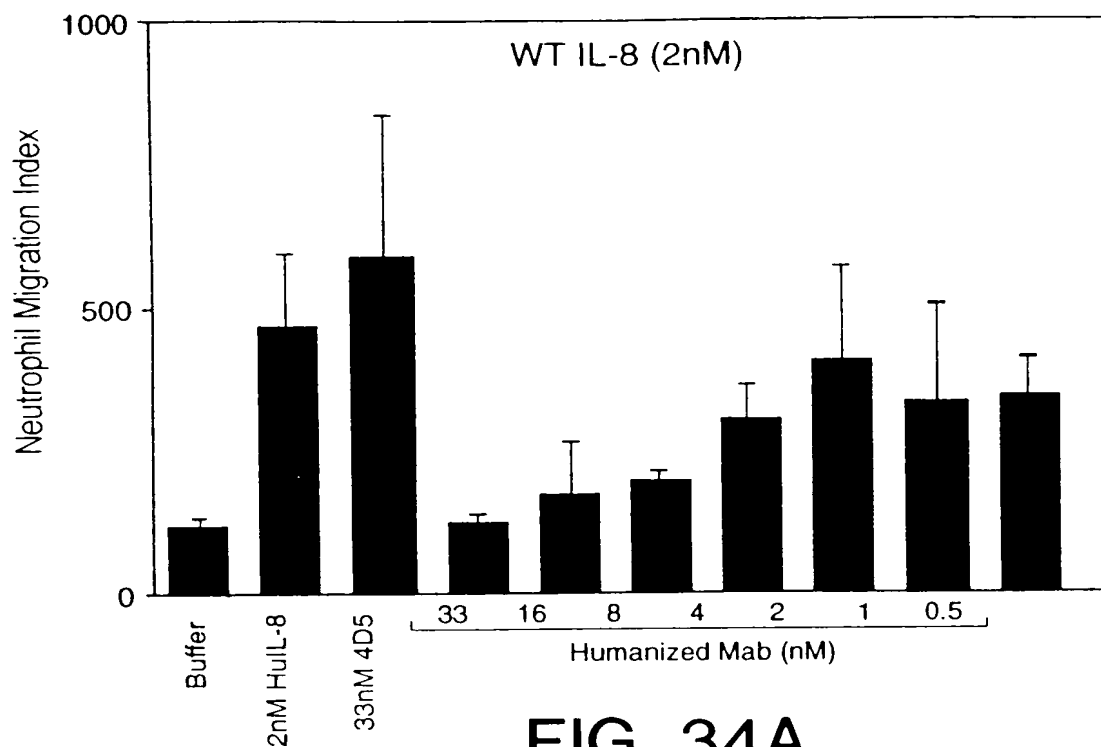


FIG. 33



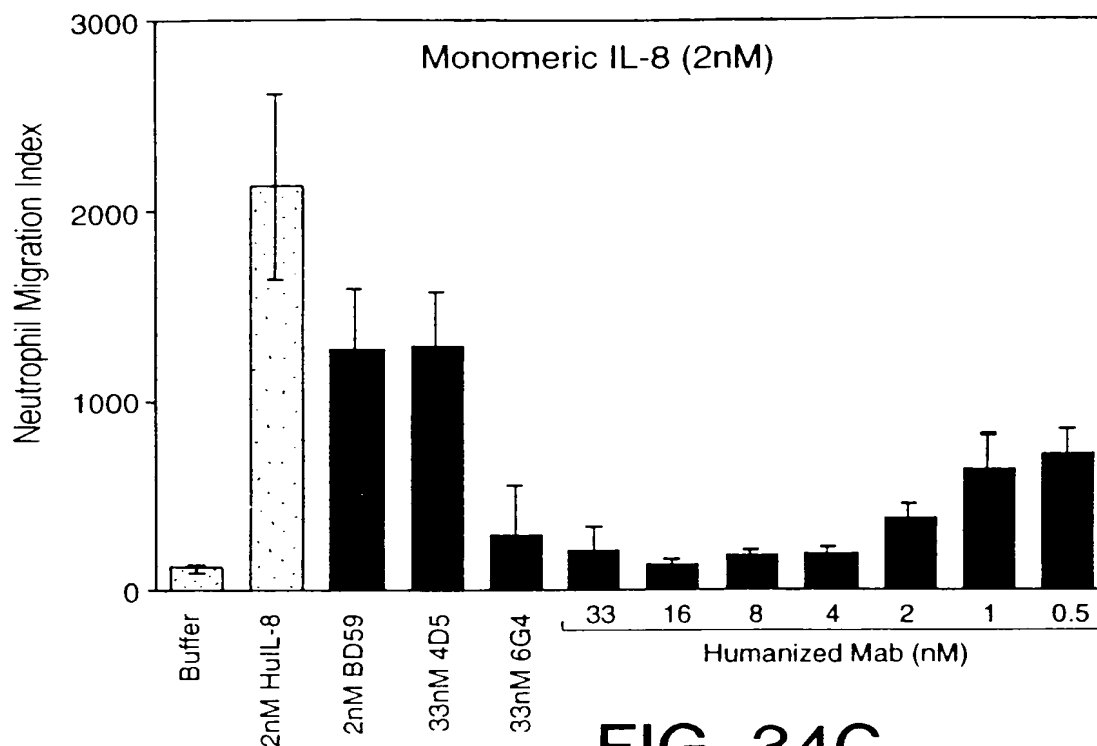


FIG. 34C

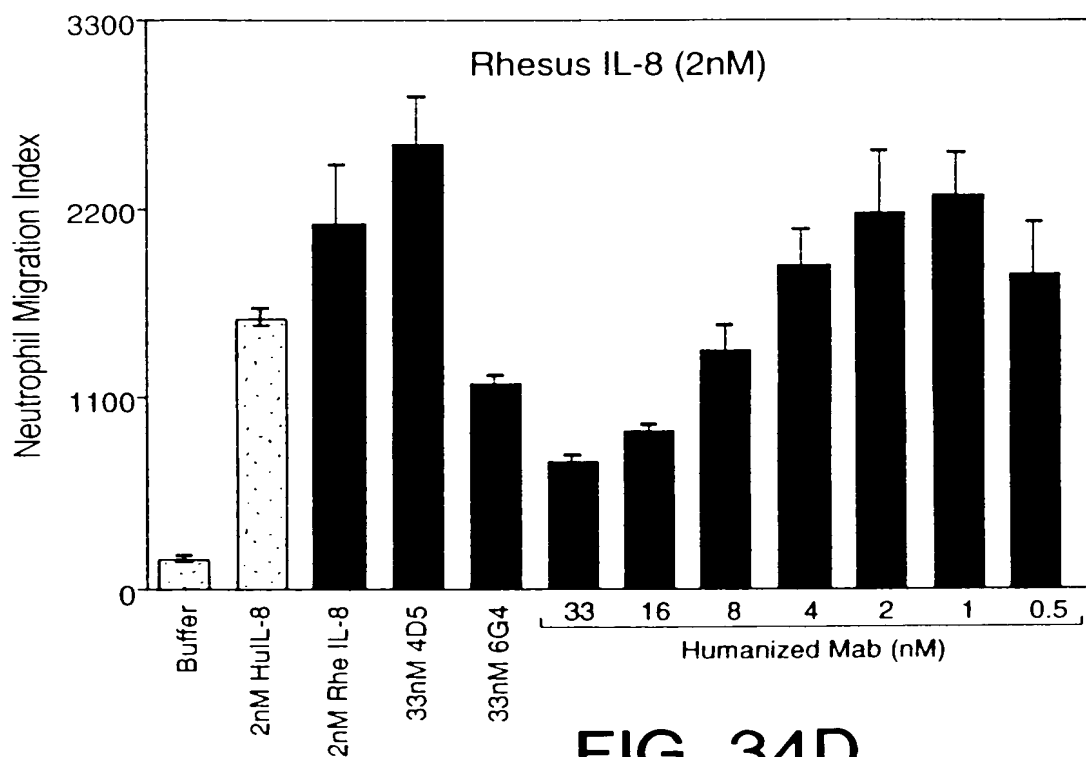


FIG. 34D

Amino Acid Sequence of the humanized anti-IL-8 6G4.2.5V11N35A Light Chain

MKKNIAFLASMFVFSIATNAYADIQMTQSPSSLSASVGDRTVITCRSSQSLVHGIGATY
LHWYQQKPGKAPKLLIYKVSNRFSGVPSRFSGSGTDFTLTISSLQPEDFATYYCSQST
HVPLTFGQGTKEIKRTVAAPSVFIFPPSDEQLKSGTASVVCLLNNFYPREAKVQWKVDN
ALQSGNSQESVTEQDSKDSYSTLSSTLTLSKADYEKHKVYACEVTHQGLSSPVTKSFNRG
EC (SEQ ID NO.56)

Amino Acid Sequence of the humanized anti-IL-8 6G4.2.5V11N35A Heavy Chain

MKKNIAFLASMFVFSIATNAYAEVQLVQSGGGLVQPGGSLRLSCAASGYSFSSHYMH
WVRQAPGKGLWVGYYIDPSNGETTYNQKFKGRFTLSRDNSKNTAYLQMNSLRAEDTAVYY
CARGDYRYNGDWFFDVWGQGTLVTVSSASTKGPSVFPLAPSSKSTSGGTAALGCLVKDYF
PEPVTVSWNSGALTSGVHTFPAVLQSSGLYSLSSVTVTPSSSLGTQTYICNVNHHKPSNTK
VDKKVEPKSCDKTHT (SEQ ID NO.52)

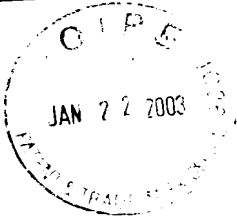
Amino Acid Sequence of the putative Pepsin Cleavage Site and GCN4 Leucine Zipper

CPPCPAPELLGGRMKQLEDKVEELLSKNYHLENEVARLKKLVGER (SEQ ID NO.57)

FIG. 35

FIG. 36

(SEO ID NO.58)



Title: Antibody Fragment-Polymer Conjugates and Humanized Anti-IL-8

Monoclonal Antibodies; Inventor: Vanessa H. et al.

Application No.: 09 234,182 (Attorney Docket No. G-92ENT-093A)

46 141

1621 GAGGACAAGG TCGAAGAGCT ACTCTCCAAG AACTACCACC TAGAGAATGA AGTGGCAAGA
CTCCTGTTCC AGCTTCTCGA TGAGAGGTTC TTGATGGTGG ATCTCTTACT TCACCGTTCT
248 E D K V E E L L S K N Y H L E N E V A R

1681 CTCAAAAAGC TTGTCGGGGA GCGCTAA (SEQ ID NO.59)
GAGTTTTTCG AACAGCCCCT CGCGATT
268 L K K L V G E R O (SEQ ID NO.60)

FIG. 37B

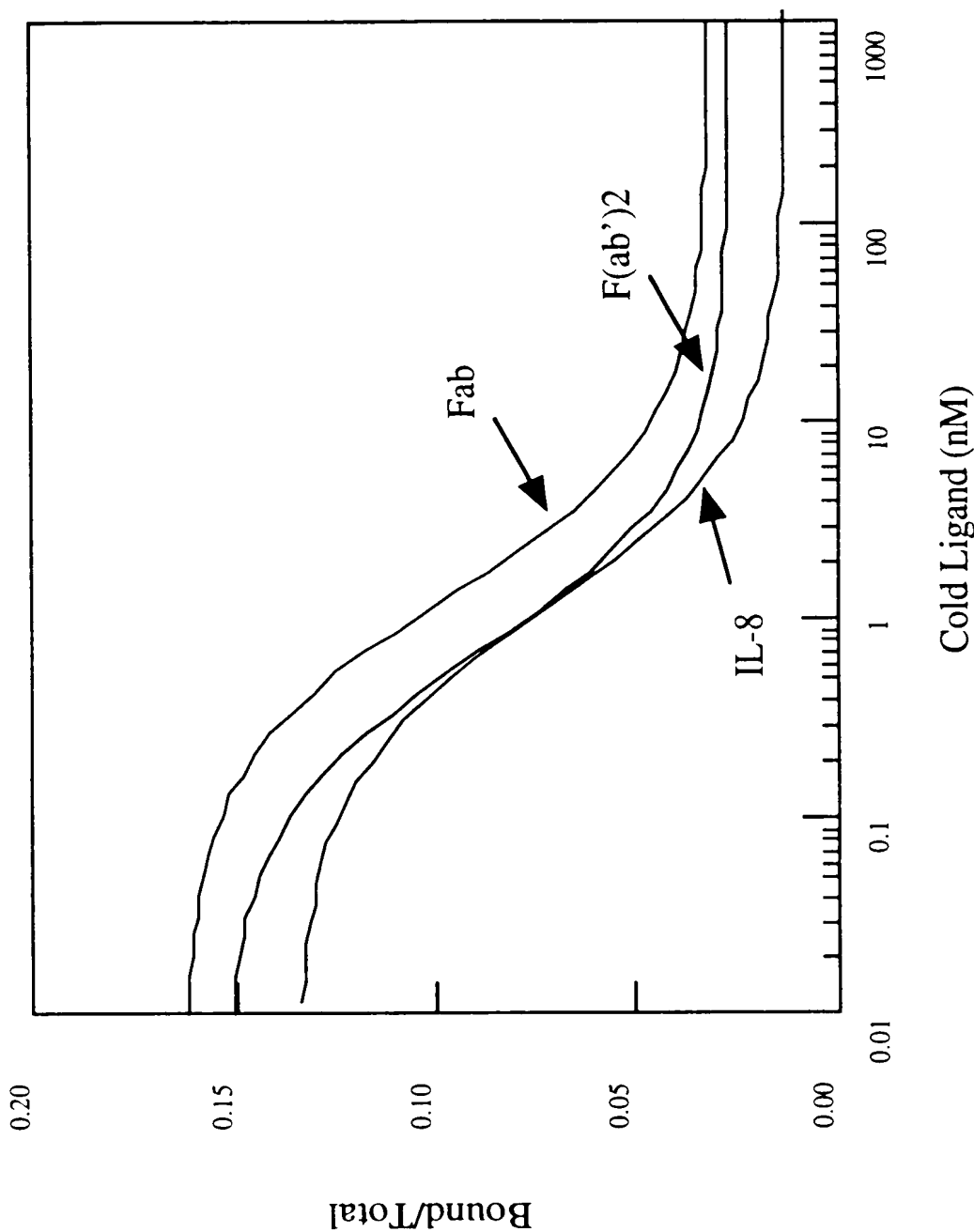


FIG. 38

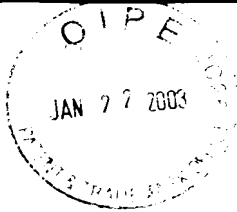
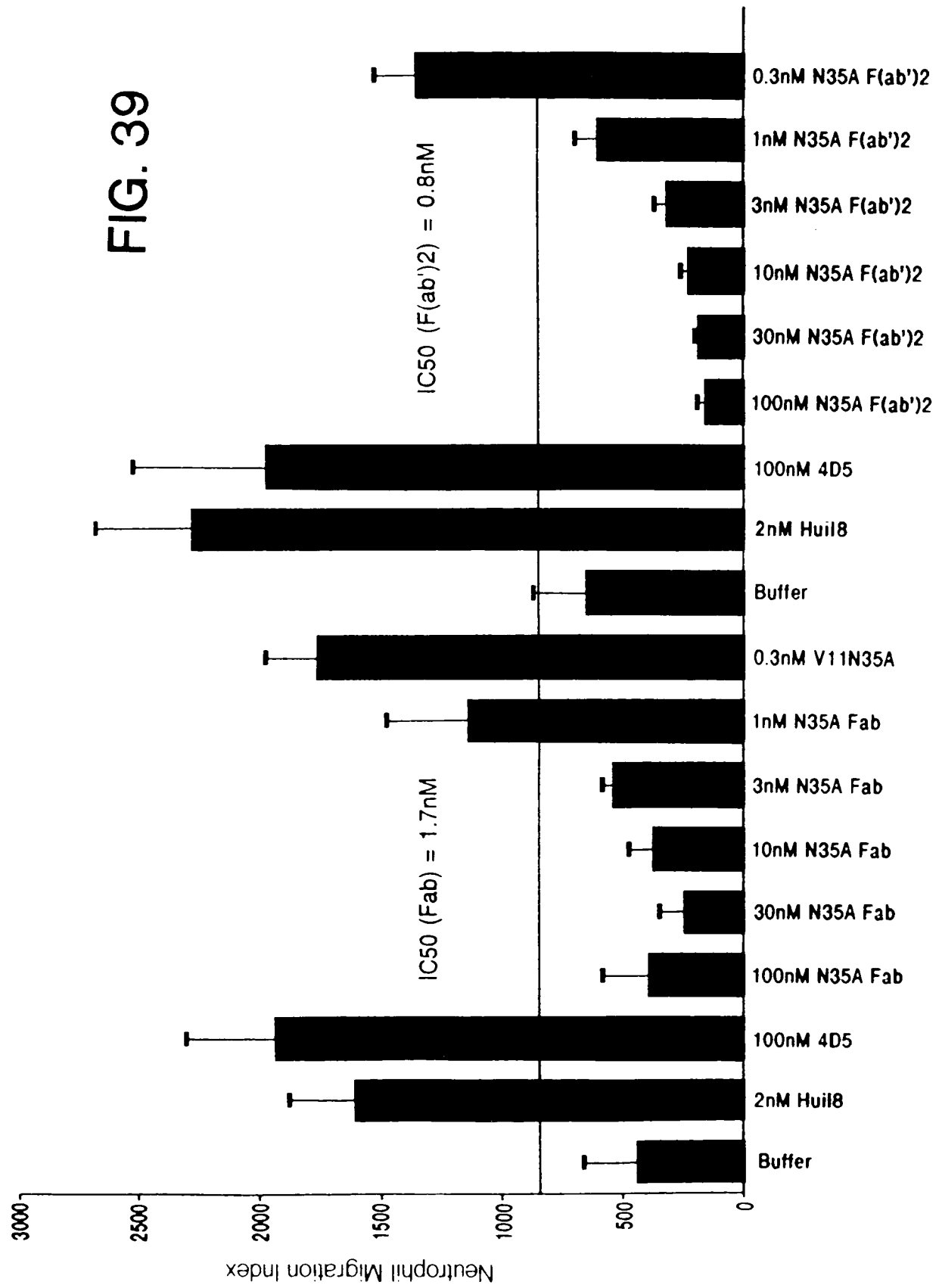
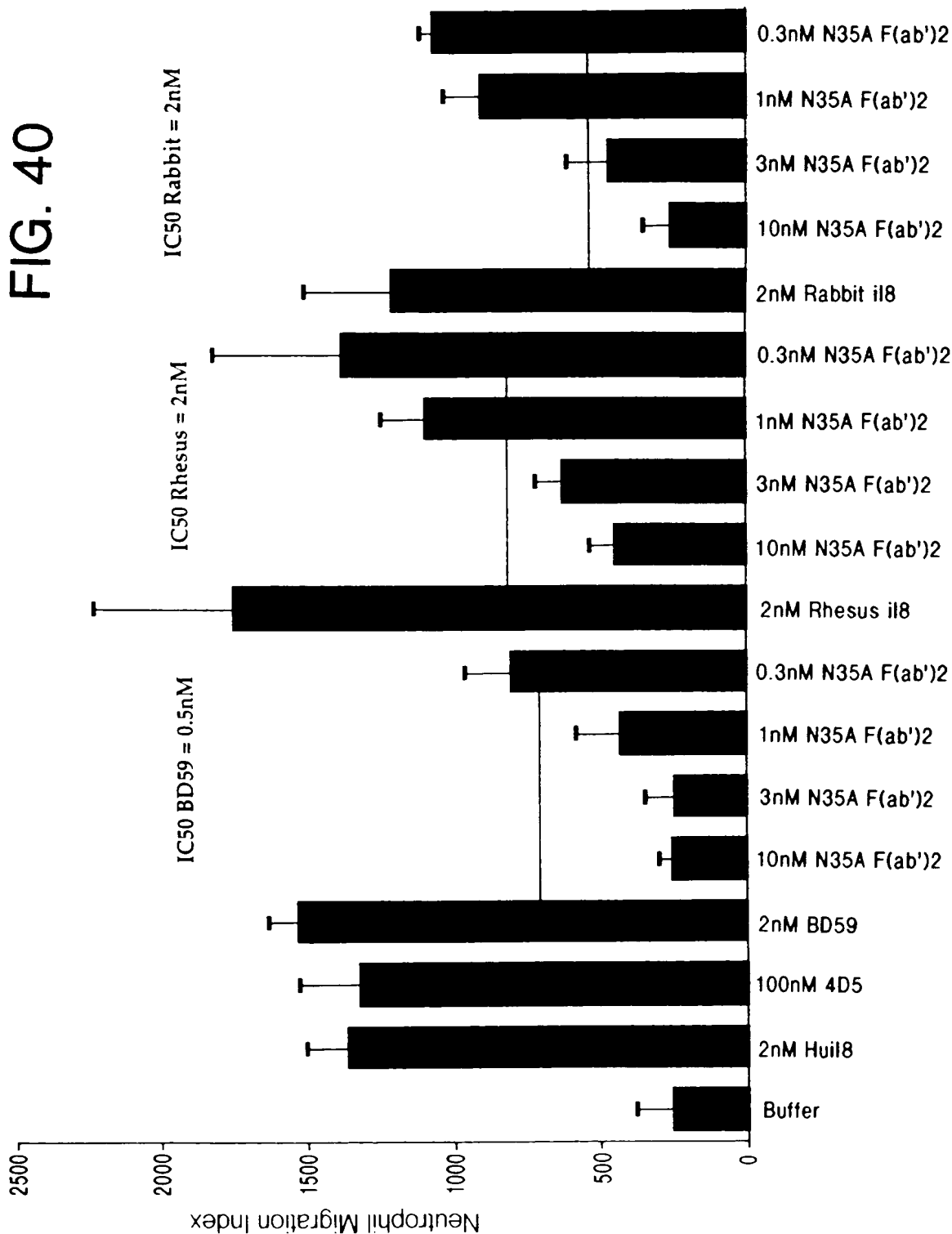


FIG. 39



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 JAN 7 2003
 100

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ecoRI      pflMI      pleI
apoI        bslI      mboII  taqI
1 GAATTCAACT TCTCCATACT TTGGATAAGG AAATACAGAC ATGAAAAATC TCATTGCTGA GTTGTATTAT AAGCTTGCC AAAAAAGAGA AGAGTCGAAT
CTTAAGTTGA AGAGGTATGA AACCTATTCC TTTATGCTG TACTTTTATG AGTAACGACT CAACAATAAA TTCGAACGGG TTTTCTCTCT TCTCAGCTTA

          bspMI      sau3AI
          hinPI      mboI/ndeII[dam-]
          hhaI/cfoI  dpnI[dam+]
          mstI      dpnII[dam-]
          avII/fspI  hinPI      aciI
          hhaI/cfoI  maeIII bsrDI  hhaI/cfoI  nspBII  bclI[dam-] mnlI
101 GAACTGTGTG CGCAGGTAGA AGCTTTGGAG ATTATCGTCA CTGCAATGCT TCGCAATATG GCGCAAAATG ACCAACAGCG GTTGATTGAT CAGGTAGAGG
CTTGACACAC GCGTCCATCT TCGAAACCTC TAATAGCAGT GACGTTACGA AGCGTTATAC CGGTTTTTAC TGGTTGTGCG CAACATACTA GTCCATCTCC

          rsaI      thal
          hinPI      fnuDII/mvnI
          hhaI/cfoI  fnu4HI
          haeII csp6I  bsoFI
          hhaI/cfoI  bbvI      maeII
          hhaI/cfoI  fnu4HI bstUI snaBI
          hhaI/cfoI  bsoFI bsh1236I mnlI
          hhaI/cfoI  bbvI hinPI bsaAI foki
          hhaI/cfoI  aluI hhaI/cfoI sfanI

          haeIII/palI
          mcrI
          eagI/xmaIII/ecI XI
          eaeI
          cfrI
          bsiEI  ahdI/eam1105I
          maeIII bsmAI
          aluI      tru9I      mseI
          pvuII      maeII  apoI  banII
          nspBII      maeI      bfaI  taqI
          tru9I      maeI      bmyI
          mseI      rmaI      bsiHKAI
          maeI      bfaI  taqI
          maeII  apoI  banII
301 AAAAGTTAAT CTTTTCAACA GCTGTCATAA AGTTGTCAG CCGGAGACTT ATAGTCGCTT TGTGTTTATT TTTTAATGTA TTTGTAACTA GAAATCGAGC
TTTTCAATTA GAAAAGTTGT CGACAGTATT TCAACAGTGC CGGCTCTGAA TATCAGCGAA ACAAAAATAA AAAATTACAT AAACATTGAT CTTAAGCTCG

```

FIG. 41A

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FIG. 41B

JAN 22 2003

Title: Antibody Fragment-Polymer Conjugates and Humanized Anti-IL-8

Monoclonal Antibodies; Inventor: Vanessa H. et al.

Application No.: 09 234,182 (Attorney Docket No. GLENENT 093A)

52 141

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scrFI      mval      ecorII      dsav      bstNI      aluI      apyl[dcn+]      bsrI
mval      ecorII      dsav      bstNI      aluI      apyl[dcn+]      bsrI
601 GGTATAGGTG CTACGTATTT ACACGTGGTAT CAACAGAAAC CAGGAAAGC TCCGAAACTA CTGATTATCA AAGTATCCAA TCGATTCTCT GGAGTCCCTT
CCATATCCAC GATCATATAA TGTGACCATG TGTGACCATG TGTGACCATG TGTGACCATG TGTGACCATG TGTGACCATG TGTGACCATG TGTGACCATG
32 G I G A T Y L H W Y Q Q K P G K A P K L L I Y K V S N R F S G V P S

mspI      hpaII      bslI      bsaWI      sau3AI      mboI/ndeII[dam-]      dpnI[dam+]      dpnII[dam-]      alwI[dam-]      nlaIV      bstYI/xhoII      bamHI
mspI      hpaII      bslI      bsaWI      sau3AI      mboI/ndeII[dam-]      dpnI[dam+]      dpnII[dam-]      alwI[dam-]      nlaIV      bstYI/xhoII      bamHI
601 CTCGTTCTC TGATCCGGT TCTGGGACGG ATTTCACTCT GACATCAGC AGTCTGCAGC CAGAAGACTT CGCAACTTAT TACTGTTCAC AGAGTACTCA
GAGCGAAGAG ACCTAGGCCA AGACCTGCG TAAAGTGAGA CTGGTAGTCG TCAGACGTCG GTCTTCTGAA GCGTTGAATA ATGACAAGTG TCTCATGAGT
66 R F S G S G S G T D F T L T I S S L Q P E D F A T Y Y C S Q S T H

styI      bsaJI      rsaI      csp6I      nlaIV      kpnI      hgiCI      banI      asp718      acc65I      maeII      bsrBI      aciI      bsmFI
styI      bsaJI      rsaI      csp6I      nlaIV      kpnI      hgiCI      banI      asp718      acc65I      maeII      bsrBI      aciI      bsmFI
801 TGTCCCGCTC ACGTTTGGAC AGGGTACCAC GGTGGAGATC AAACGAACTG TGGCTGCACC ATCTGTCTTC ATCTCCCGC CATCTGATGA GCAGTTGAAA
ACAGGGCGAG TGCACACCTG TCCCATGGTT CCACCTCTAG TTTGTTGAC ACCGACGTGG TAGACAGAG TAGACAGAG TAGACAGAG TAGACAGAG TAGACAGAG
99 V P L T F G Q G T K V E I K R T V A A P S V F I F P P S D E Q L K

tfiI      hinfi      bsmFI      taqI      bpmI/gsuI[dcn-]      claI/bsp106      pleI      bspDI[dam-]      hinfi
tfiI      hinfi      bsmFI      taqI      bpmI/gsuI[dcn-]      claI/bsp106      pleI      bspDI[dam-]      hinfi
601 GGTATAGGTG CTACGTATTT ACACGTGGTAT CAACAGAAAC CAGGAAAGC TCCGAAACTA CTGATTATCA AAGTATCCAA TCGATTCTCT GGAGTCCCTT
CCATATCCAC GATCATATAA TGTGACCATG TGTGACCATG TGTGACCATG TGTGACCATG TGTGACCATG TGTGACCATG TGTGACCATG TGTGACCATG
32 G I G A T Y L H W Y Q Q K P G K A P K L L I Y K V S N R F S G V P S

fnu4HI      bsoFI      bbvI      scfI      mboII      pstI      bpuAI      bbsI      rsaI      csp6I      scaI      nlaIII
fnu4HI      bsoFI      bbvI      scfI      mboII      pstI      bpuAI      bbsI      rsaI      csp6I      scaI      nlaIII
701 CTCGTTCTC TGATCCGGT TCTGGGACGG ATTTCACTCT GACATCAGC AGTCTGCAGC CAGAAGACTT CGCAACTTAT TACTGTTCAC AGAGTACTCA
GAGCGAAGAG ACCTAGGCCA AGACCTGCG TAAAGTGAGA CTGGTAGTCG TCAGACGTCG GTCTTCTGAA GCGTTGAATA ATGACAAGTG TCTCATGAGT
66 R F S G S G S G T D F T L T I S S L Q P E D F A T Y Y C S Q S T H

sau3AI      mboI/ndeII[dam-]      dpnI[dam+]      dpnII[dam-]      bbvI      mboII      bpuAI      bbsI      aciI
sau3AI      mboI/ndeII[dam-]      dpnI[dam+]      dpnII[dam-]      bbvI      mboII      bpuAI      bbsI      aciI
801 TGTCCCGCTC ACGTTTGGAC AGGGTACCAC GGTGGAGATC AAACGAACTG TGGCTGCACC ATCTGTCTTC ATCTCCCGC CATCTGATGA GCAGTTGAAA
ACAGGGCGAG TGCACACCTG TCCCATGGTT CCACCTCTAG TTTGTTGAC ACCGACGTGG TAGACAGAG TAGACAGAG TAGACAGAG TAGACAGAG TAGACAGAG
99 V P L T F G Q G T K V E I K R T V A A P S V F I F P P S D E Q L K

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FIG. 41C

FIG. 41D

-23

1301 CTACAAACGC GTACGGCTGAG GTTCAGCTAG TGCAGTCTGG CGGTGGCGCTG GTGCAGCCAG GTGGCTCACT CCGTTTGTCC TGTGCAGCTT CTGGCTACTC
GATGTTTGGC GATGGCACTC CAAGTCGATC ACGTCAGACC GCCACCGGAC CACGTCGGTC CCCCGAGTGA GGCAACACAGG ACACGTCGAA GACCGATGAG

scrFI
nciI
mspI
hpaII
dsaV
cauII
bslI
xmaI/pspAI
smaI
scrFI
nciI
dsaV
cauII
bslI
scrFI
mvaI
ecoRII

FIG. 41E

FIG. 41F

FIG. 41G

FIG. 41H

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FIG. 41I

FIG. 41J

FIG. 41K

[illegible]

Accession	Protein	Gene	Strain	Host	Location	Size (aa)	Size (bp)	GC (%)	RefSeq
3301	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3302	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3303	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3304	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3305	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3306	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3307	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3308	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3309	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3310	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3311	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3312	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3313	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3314	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3315	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3316	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3317	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3318	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3319	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3320	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3321	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3322	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3323	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3324	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3325	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3326	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3327	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3328	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3329	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3330	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3331	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3332	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3333	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3334	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3335	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3336	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3337	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3338	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3339	ftsH	ftsH	ftsH	ftsH	ftsH	111	333	50.15	NC_016416.1
3340	fts								

[illegible]

FIG. 41L

FIG. 41M



3801 TGGTCTTCGG TTTCGGTGT TCGTAAAGTC TGGAAACGCG GAAGTCAGCG CCTGACACCA TTATGTTCCG GATCTGCATC GCAGGATGCT GCTGGCTACC
ACCAAGAGCC AAAGGCACAA AGCAATTCAG ACCTTTGCGC CTTCACTCGC GGGACGTGGT AATACAAGGC CTAGACGTAG CGTCTACGA CGACCGATGG

3901 CTGTGGAACA CCTACATCTG TATTACGAA GCGCTGGCAT TGACCCCTGAG TGATTTTCT CTGGTCCCGC CGCATCCATA CCGCCAGTTG TTTACCCCTCA
GACACCTTGT GGATGTAGAC ATAATTGCTT CGCGACCGTA ACTGGGACTC ACTAAAAGA GACCAGGCGC GCGTAGGTAT GCGGTCAAC AAATGGAGT

4001 CAACGTTCCA GTAACCGGCG ATGTTTCATCA TCAGTAACCC GTATCGTGAG CATCCTCTCT CGTTTCATCG GTATCATTAC CCCCATGAAC AGAAATTCCTC
GTTGCAAGGT CATTGGCCCG TACAAGTACT AGTCATTGGG CATAGCACTC GTAGGAGAGA GCAAAGTAGC CATAGTAATG GGGGTACTTG TCTTTAAGGG

Restriction Enzymes and Sites:

Left side (3801):

- sau3AI
- mboI/ndeII(dam-)
- mamI(dam-)
- dpnI(dam+)
- dpnII(dam-)
- bstYI/xhoII
- alwI(dam-)
- mspI
- hpaII
- mroI/bsaBI(dam-)
- bspMI
- bspEI(dam-)
- bsaWI
- sfanI
- accIII(dam-)
- fokI
- cac8I
- fnu4HI
- bsaFI
- bbvI

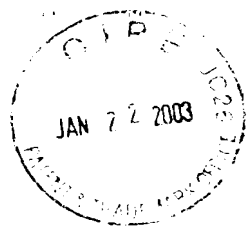
Right side (3901):

- aciI
- bsmFI
- fokI
- sau96I
- sfanI
- nlaIV
- aciI
- avaII
- fnu4HI
- bsrI
- asuI
- bsaFI
- aciI
- mnII

Bottom (4001):

- nspl
- scrFI
- ncil
- mspI
- hpaII
- bsrI
- bslI
- maeII
- psp1406I
- maeIII
- nspHI
- cauII
- dsav
- nlaIII
- maeIII
- sfanI
- mnII
- fokI
- nlaIII
- apoI
- bslI

FIG. 41N



cac8I
 sau96I
 tru9I haeIII/palI
 mseI asuI
 mnlI maelII
 aciI bslI nlaIII aciI
 bpmI/gsuI[dcM-]
 4101 CCTTACACGG AGGCATCAAG TGACCAACA GGAAGAAACC GCCCTTAACA TGCCCGCTT TATCAGAAGC CAGACATTAA CGTCTCTGGA GAAACTCAAC
 GGAATGTGCC TCCGTAGTTC ACTGGTTTGT CCTTTTGG CGGGAATTGT ACCGGCCGAA ATAGTCTTCG GTCTGTAAAT CGGAAGACCT CTTTGAGTTG
 fnu4HI
 bsoFI fnuDII/mvnI
 aluI bstUI
 pvuII hinPI
 nspBII hhaI/cfoI
 fnu4HI
 bsoFI fnuDII/mvnI
 bcgI bstUI
 bbvI mnlI bsh1236I
 aciI hgaI foki
 fnuDII/mvnI
 bstUI
 bsh1236I
 aluI hgaI foki
 4201 GAGCTGGACG CGGATGAACA GGCAGACATC TGTGAATCGC TTCACGACCA CGCTGATGAG CTTTACCGCA GCTGCCCTCG CGGTTTCGGT GATGACGGTG
 CTCGACCTGC GCCTACTTGT CCGTCTGTAG ACACCTAGCG AAGTGCTGGT GCGACTACTC GAAATGGCGT CGACGGAGCG CGCAAAGCCA CTACTGCCAC
 esp3I
 bsmBI
 bsmAI
 mspI
 fnu4HI hpaII
 bsoFI scrFI
 bbvI nciI
 nlaIII dsav
 nspI cauII
 nspHI aluI bslI
 mnlI
 4301 AAAACCTCTG ACACATGCAG CTCCCGGAGA CGGTACAGC TTGTCTGTAA CGGATGCCG GGAGCAGACA AGCCCGTCAG GCGCGTCAG CGGGTGTGG
 TTTTGGAGAC TGTGTACGTC GAGGGCCTCT GCCAGTGTG GCAAGACATT CGCCTACGGC CCTCGTCTGT TCGGGCAGTC CCGCGCAGTC GCCCACAACC
 hgaI
 thaI
 fnuDII/mvnI
 bstUI aciI
 bsh1236I
 hinPI nspBII
 hhaI/cfoI
 aciI
 scrFI
 nciI
 mspI
 hpaII
 sfaNI
 foki dsav
 aciI cauII
 drdI
 fnuDII/mvnI
 bstUI aciI
 bsh1236I
 hinPI nspBII
 hhaI/cfoI
 aciI

FIG. 410

mspI
 hpaII
 sau3AI
 mboI/ndeII[dam-]
 dpnI[dam+]
 dpnII[dam-]
 aluI
 aluI/dam-]
 aluI/dam-]
 nspBII
 aciI
 aciI
 fnu4HI
 bsoFI
 bbvI
 cac8I
 TTTTGTGTTG CAAGCAGCAG
 GTTCGTGCTC

maeIII
 eco57I bsrI
 TCGGAAAAAG AGTTGTGAGC TCCTGATCCG GCAAAACAAAC CACCGCTGGT AGCGGTGGT TTTTGTGTTG CAAGCAGCAG
 AGACGACTTC GGTCAATGGA AGCCTTTTTC TCAACCATCG AGAAGTGGC CGTTGTTTG GTGGCAGCA TCGCCACCA AAAAACCAAC GTTCGTGCTC

sau3AI
 mboI/ndeII[dam-]
 mboI/ndeII[dam-]
 sau3AI mboI/ndeII[dam-]
 mboI/ndeII[dam-]
 sau3AI mboI/ndeII[dam-]
 mboI/ndeII[dam-]
 dpnI[dam+]
 dpnII[dam-]
 dpnI[dam+]
 dpnII[dam-]
 bstYI/xhoII aluI/dam-]
 bstYI/xhoII aluI/dam-]
 aluI/dam-]
 bstYI/xhoII
 bsh1236I
 ATCTCAAGAA GATCCTTTGA TCCTTTCTAC GGGGTCTGAC GCTCAGTGA ACAGAACTC ACGTTAAGG ATTTTGGTCA
 TAATGCGCGT CTTTTTTC TAGAGTTCTT CTAGGAACT AGAAAGATG CCCCAGACTG CGAGTCACCT TGCTTTTGAG TGAATTCCC TAAACACAGT

sau3AI
 mboI/ndeII[dam-]
 rmaI
 hphI dpnI[dam+]
 mboII[dam-]
 sau3AI maeI tru9I
 mboI/ndeII[dam-] msei
 dpnI[dam+]
 dpnII[dam-]
 dpnI[dam-]
 aluI/dam-]
 bstYI/xhoII bstYI/xhoII msei
 aluI/dam-]
 bfaI ahaIII/draI
 ahaIII/draI
 tru9I msei
 maeIII
 TGAAGTATC AAAAGGATC TTCACCTAGA TCCTTTTAAA TTAATAATGA AGTTTAAAT CAATCTAAAG TATATATGAG TAAACTTGGT CTGACAGTTA
 ACTCTAATAG TTTTTCCTAG AAGTGGATCT AGGAAATTT AATTTTACT TCAAAATTTA GTTAGATTC ATAPATACTC ATTTGAACCA GACTGTCAAT

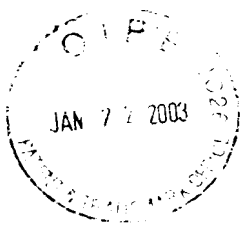
nlaIV
 hgiCI
 banI
 mnlI
 sau3AI
 mboI/ndeII[dam-]
 dpnI[dam+]
 dpnII[dam-]
 ddel
 foki
 ahdI/eam1105I
 pleI
 hinfI
 mnlI
 CCAATGCTTA ATCAGTGAGG CACCTATCTC AGCGATCTGT CTATTTCTGT CATCCATAGT TGCCTGACTC CCGCTCGTGT AGATAACTAC GATACGGGAG
 GGTACCAAT TACTCACTCC GTGGATAGAG TCGCTAGACA GATAAGCAA GTAGGTATCA ACGGACTGAG GGCAGCACA TCTATTGATG CTATGCCCTC

FIG. 41R

FIG. 41S

6001 ATTCTCTTAC TGTCATGCCA TCCGTAAGAT GCTTTTCTGT GACTGGTGAG TACTCAACCA ACTCATTCTG AGAATAGTGT ATGCGCGGAC CGAGTTGCTC
 TAAGAGAAATG ACAGTACGGT AGGCATTCTA CGAAAGACA CTGACCACTC ATGAGTTGGT TCAGTAAGAC TCTTATCACA TACGCGGCTG GCTCAACGAG
 mcrI
 bsiEI
 bcgI
 fnu4HI
 bsoFI
 acII
 rsal
 bsrI
 scaI
 maeIII hphI csp6I
 ddel
 hgaI
 hinPI
 hhaI/cfoI
 mspI
 hpaII
 scrFI
 nciI
 dsav
 cauII hincII/hindII
 aciI
 hnuDII/mvnI
 bstUI
 bsh1236I
 hgiAI/aspHI
 bsp1286
 tru9I bsiHKAI
 mseI bmyI
 ahaII/draI
 asp700 mboII
 sau3AI
 mboI/ndeII[dam-]
 dpnI[dam+]
 dpnII[dam-]
 bstYI/xhoII
 alwI[dam-]
 6101 TTGCCCCGGC TCAACACGGG ATAATACCGC GCACATAGC AGAATCTTAA AAGTGCTCAT CATTTGGAAA CGTTCTTCGG GCGGAAACT CTCRAGGATC
 AACGGCCGC AGTTGTGCC TATTATGGC CGGTGTATCG TCTTGAAATT TTCACGAGTA GTACCTTTT GCAAGAAGCC CCGCTTTTGA GAGTTCTCTAG
 hgiAI/aspHI
 bsrI
 sau3AI
 taqI
 mboI/ndeII[dam-]
 dpnI[dam+]
 dpnII[dam-]
 alwI[dam-]
 bstYI/xhoII
 maeIII
 bssSI
 hphI
 6201 TTACCGCTGT TCAGATCCAG TTCGATGTAA CCACTCGTG CACCCAACTG ATCTTCAGCA TCTTTTACTT TCACCCAGCGT TTCTGGGTGA GCAAAAACAG
 AATGGGACA ACTCTAGGTC AAGCTACATT GGGTGAGCAC GTGGGTTGAC TAGAAGTCGT AGAAATGAA AGTGGTCGCA AAGACCCACT CGTTTTTGTC
 aciI
 fnu4HI
 bsoFI
 mboII
 earI/ksp632I
 sspI
 6301 GAAGGCAAAA TGCCGCAAAA AAGGGAATAA GGGGACACG GAATGTTGA ATACTCATAC TCTTCCTTTT TCAATATTAT TGAAGCATTT ATCAGGTTTA
 CTTCCGTTTT ACGGCGTTTT TTCCCTTATT CCGGCTGTGC CTTTACAAC TATGAGTATG AGAAGGAAAA AGTATAATA ACTTCGTAAA TAGTCCCAAT

FIG. 41T



APPROVED	O.G. FIG.	
BY	CLASS	SUBCLASS
DRAFTSMAN		

```

        hinPI
        thal
        fnuDII/mvni
        bstUI
        bsh1236I
        aciI
        nlaIV hhaI/cfoI
        nlaIII
        rcaI
        bspHI aciI
        bsmAI bsrBI
        6401 TTCTCTCATG AGCGGATACA TATTGAATG TATTAGAAA AATAACAAA TAGGGTTCC GCGCACATTT CCGCGAAAAG TGCCACCTGA CGTCTAAGAA
        AACAGAGTAC TCGCCTATGT ATAACTTAC ATAAATCTTT TTATTGTTT ATCCCCAAGG CGCGGTAAA GGGGCTTTTC ACGGTGGACT GCAGATTCTT
        sau96I
        haeIII/palI
        asuI
        ecoO109I/draII
        mnlI
        bssSI
        bpuAI
        bbsI
        nlaIII
        rcaI
        bspHI
        6501 ACCATTATTA TCATGACATT AACCTATAAA AATAGGCGTA TCACGAGGCC CTTTCGTCTT CAA (SEQ ID NO 61)
        TGGTAATAAT AGTACTGTAA TTGGATATTT TTATCCGCAT AGTGCTCCGG GAAAGCAGAA GTT
```

FIG. 41U

JAN 27 2003

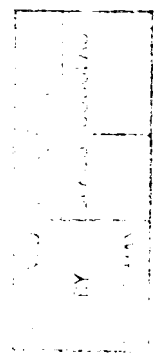
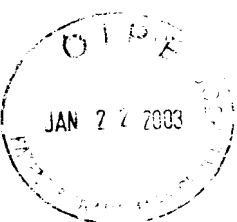
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```

aatII(GACGTC): 1645 6489
acc65I(GGTACC): 403 823
accI(GTMKAC): 1093 1963 4449
accII(TCCGGA): 3867[dam-]
accI(CCGC): 178 542 805 877 1340 1750 1826 2011 2039 2043 2182 2242 2384 2492 2501 2504
2628 2781 2784 2787 2906 2926 3005 3045 3094 3141 3226 3241 3309 3342 3367 3412
3436 3448 3490 3544 3597 3613 3619 3700 3838 3967 3970 3981 4139 4155 4210 4266
4351 4390 4400 4442 4467 4505 4518 4544 4561 4604 4611 4632 4723 4751 4878 4897
5018 5128 5263 5272 5634 5725 5916 5962 6083 6127 6204 6313 6412 6459
see hinI
acyI 1307 4678
afIIII(ACRYGT): 1788
ageI(ACCGGT): 1645 1813 2616 2637 2751 3408 6107 6489
ahaII/bsaII(GRCGYC): 5435 5454 6146
ahaII/draI(TTTAAA): 72 121 252 320 398 532 589 648 1126 1144 1167 1325 1386 1906 2054 2075 2126
ahdI/eamI105I(GACNNNNNGTC): 346 5566
aluI(AGCT): 2218 2233 2889 3292 4202 4259 4270 4319 4338 4619 4845 4935 4981 5238 5759 5859
5922
alw44I/snoI(GTGCAC): 1831 4494 4992 6238
alwI(dam-)(GGATC): 412 413 712 713 1171 1471 2578 2579 3300 3870 5245 5319 5331 5416 5429 5893
6196 6214
alwNI(dcm-)(CAGNNNCTG): 1117 1385 5089
apaI(GGGCCC): 1695
apaLI/snoI(GTGCAC): 1831 4494 4992 6238
apoI(RAATTY): 1 391 4093
apyI(dcm+)(CCWGG): 640 999 1347 1357 1449 1665 1713 1755 1764 2333 3262 3645 4705 4826 4839
aseI/asnI/vspI(ATTAAAT): 5742
asni see aseI
asp700(GAANNNTTC): 905 930 4234 6166
asp718(GGTACC): 403 823
asphi see hgiAI
aspi see tthlII
asuI(GGNCC): 1119 1195 1425 1434 1446 1512 1695 1696 1752 2155 2375 2727 3002 3090 3339 3463

```

FIG. 41V



Stop Template Primer

SL.97.2 5' CAT GGT ATA GGT TAA ACT TAT TTA CAC 3' (SEQ ID NO.63)

NNS Randomization Primer

SL.97.3 5' CAT GGT ATA GGT NNS ACT TAT TTA CAC 3' (SEQ ID NO.64)

FIG. 42



Randomization of Position N35 of Variable Light Chain CDR-1 Amino Acid Frequency

Phage Display (NNS Codon Library) Sort #3

Amino Acid	Frequency	% Total	IC50 (nM)
Asparagine (wt)	1	5.6	4.9
Glycine	6	16.6	3.1
Aspartic Acid	3	16.6	3.1
Glutamic Acid	4	22.2	0.1
Alanine	2	5.6	0.2
Lysine	1	5.6	ND
Serine	1	1.9	ND

FIG. 43A

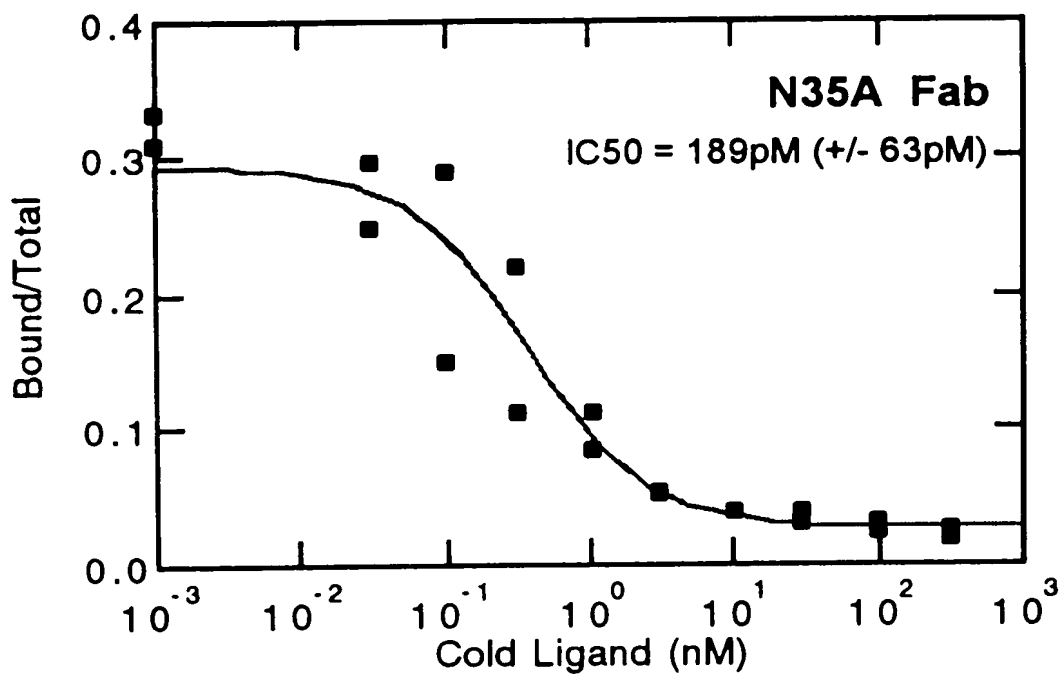
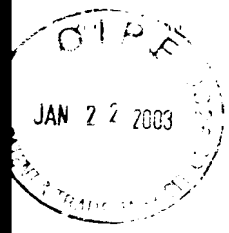


FIG. 43B

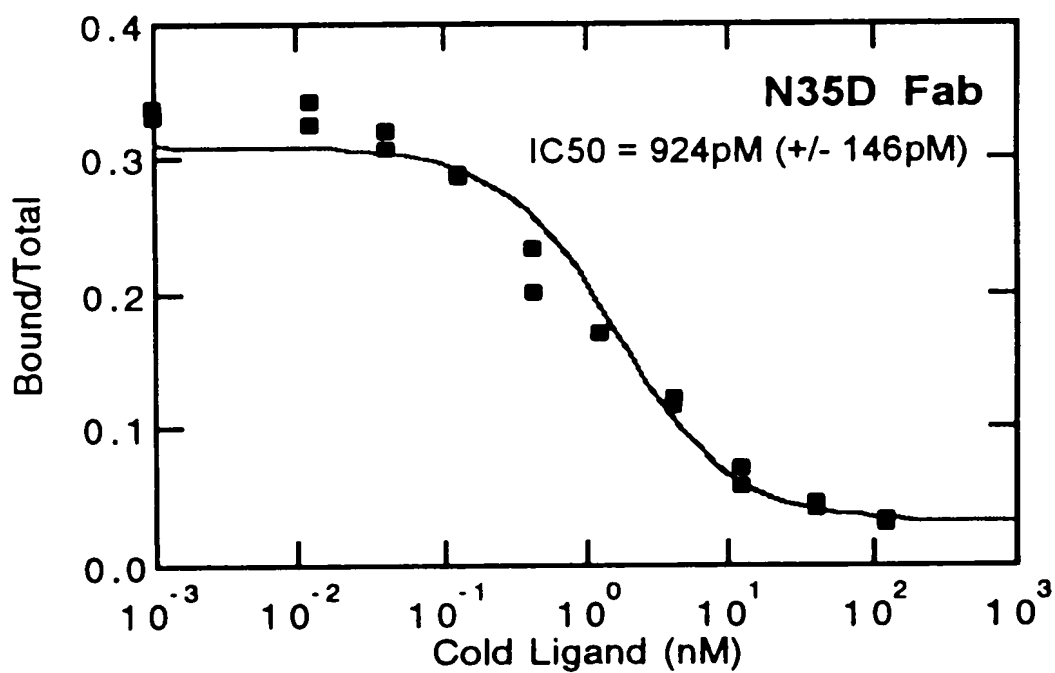


FIG. 43C

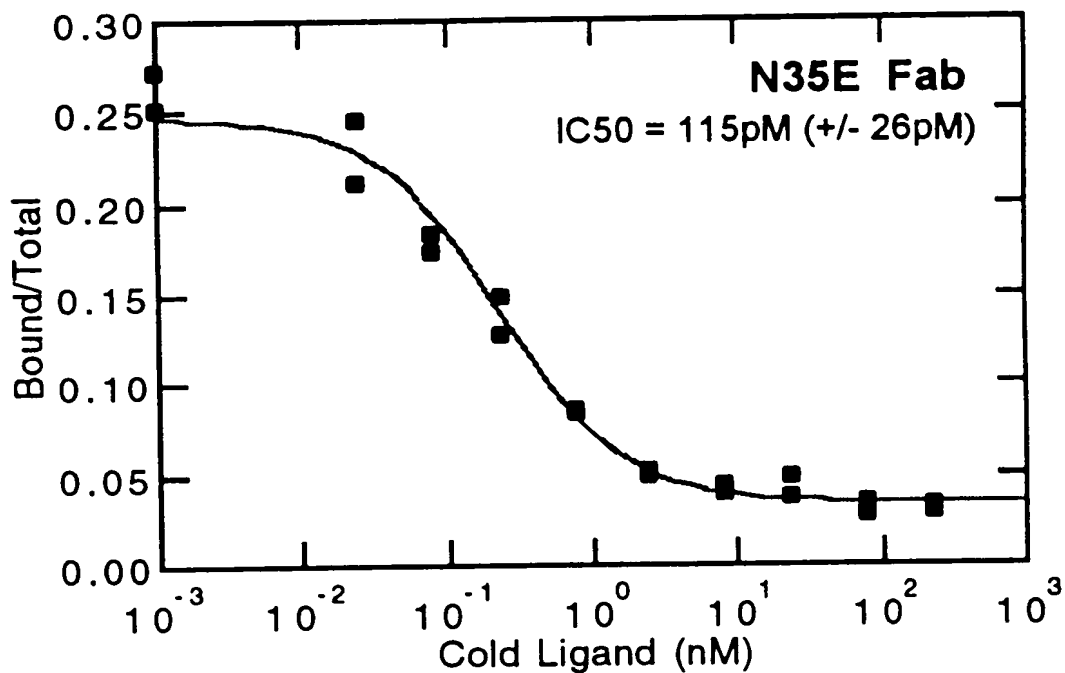


FIG. 43D

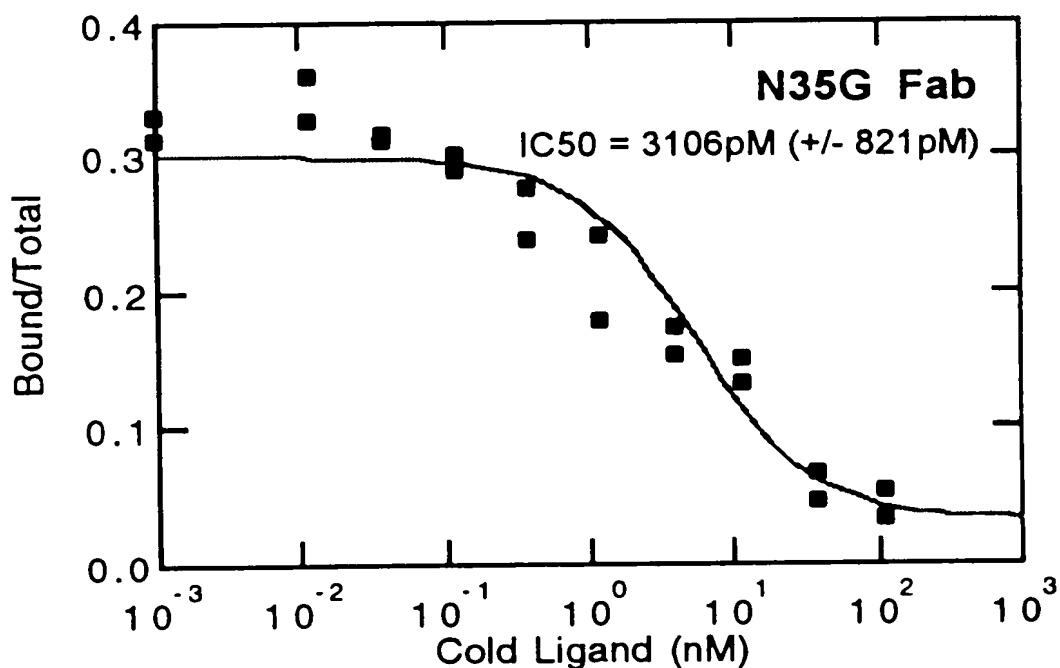
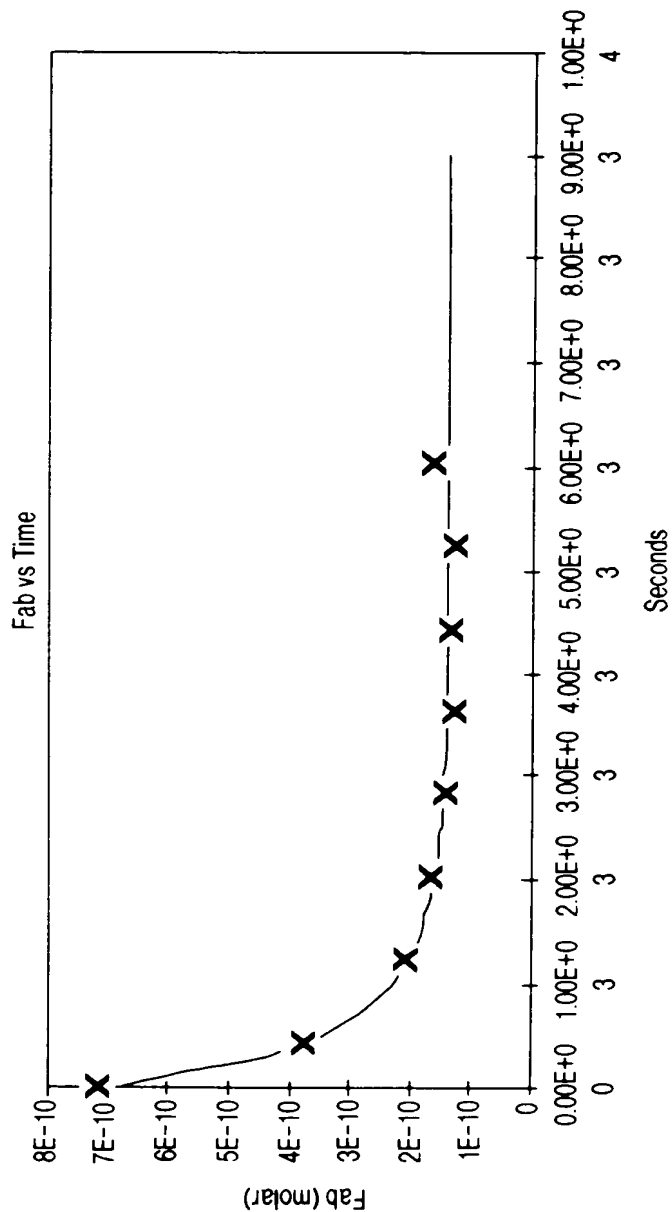
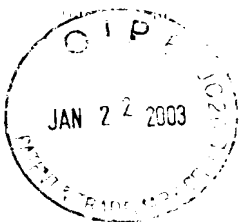


FIG. 43E



Representative Conc versus Time Plot. Shown is the kinetic data for 6G4V11N35A.F(ab')₂.

SAMPLE	ka	kd	Kd
6G4V11N35A-Fab	ND	ND	114pM
6G4V11N35A-F(ab') ₂	2.0x10 ⁶	2.1x10 ⁻⁴	109pM
6G4V11N35E-Fab	4.7x10 ⁶	2.6x10 ⁻⁴	54pM

FIG. 44



1 ATGAAAAAGA ATATCGCATT TCTTCTTGCA TCTATGTTTCG TTTTCTCTAT TGCTACAAAC
TACTTTTCT TATAGCGTAA AGAAGAACGT AGATACAAGC AAAAAAGATA ACGATGTTTG
-23 M K K N I A F L L A S M F V F S I A T N

61 GCATACGCTG ATATCCAGAT GACCCAGTCC CCGAGCTCCC TGTCCGCCTC TGTGGGCGAT
CGTATGCGAC TATAGGTCTA CTGGGTCAGG GGCTCGAGGG ACAGGCGGAG ACACCCGCTA
-3 A Y A D I Q M T Q S P S S L S A S V G D

121 AGGGTCACCA TCACCTGCAG GTCAAGTCAA AGCTTAGTAC ATGGTATAGG TGAGACGTAT
TCCAGTGGT AGTGGACGTC CAGTTCAGTT TCGAATCATG TACCATATCC ACTCTGCATA
18 R V T I T C R S S Q S L V H G I G E T Y

181 TTACACTGGT ATCAACAGAA ACCAGGAAAA GCTCCGAAAC TACTGATTTA CAAAGTATCC
AATGTGACCA TAGTTGTCTT TGGTCCTTTT CGAGGCTTTG ATGACTAAAT GTTTCATAGG
38 L H W Y Q Q K P G K A P K L L I Y K V S

241 AATCGATTCT CTGGAGTCCC TTCTCGCTTC TCTGGATCCG GTTCTGGGAC GGATTTCACT
TTAGCTAAGA GACCTCAGGG AAGAGCGAAG AGACCTAGGC CAAGACCCTG CCTAAAGTGA
58 N R F S G V P S R F S G S G S G T D F T

301 CTGACCATCA GCAGTCTGCA GCCAGAAGAC TTCGCAACTT ATTACTGTTC ACAGAGTACT
GACTGGTAGT CGTCAGACGT CGGTCTTCTG AAGCGTTGAA TAATGACAAG TGTCTCATGA
78 L T I S S L Q P E D F A T Y Y C S Q S T

361 CATGTCCCGC TCACGTTTGG ACAGGGTACC AAGGTGGAGA TCAAACGAAC TGTGGCTGCA
GTACAGGGCG AGTGCAAACC TGTCCCATGG TTCCACCTCT AGTTTGCTTG ACACCGACGT
98 H V P L T F G Q G T K V E I K R T V A A

421 CCATCTGTCT TCATCTTCCC GCCATCTGAT GAGCAGTTGA AATCTGGAAC TGCTTCTGTT
GGTAGACAGA AGTAGAAGGG CGGTAGACTA CTCGTCAACT TTAGACCTTG ACGAAGACAA
118 P S V F I F P P S D E Q L K S G T A S V

481 GTGTGCCTGC TGAATAACTT CTATCCCAGA GAGGCCAAAG TACAGTGGAA GGTGGATAAC
CACACGGACG ACTTATTGAA GATAGGGTCT CTCCGGTTTC ATGTCACCTT CCACCTATTG
138 V C L L N N F Y P R E A K V Q W K V D N

541 GCCCTCCAAT CGGGTAACTC CCAGGAGAGT GTCACAGAGC AGGACAGCAA GGACAGCACC
CGGGAGGTTA GCGCATTTGAG GGTCTCTCTA CAGTGTCTCG TCCTGTCGTT CCTGTCTGTG
158 A L Q S G N S Q E S V T E Q D S K D S T

601 TACAGCCTCA GCAGCACCTT GACGCTGAGC AAAGCAGACT ACGAGAAACA CAAAGTCTAC
ATGTCGGAGT CGTCGTGGGA CTGCGACTCG TTTCGTCTGA TGCTCTTTGT GTTTCAGATG
178 Y S L S S T L T L S K A D Y E K H K V Y

661 GCCTGCGAAG TCACCCATCA GGGCCTGAGC TCGCCCGTCA CAAAGAGCTT CAACAGGGGA
CGGACGCTTC AGTGGGTAGT CCCGGACTCG AGCGGGCAGT GTTCTCTGAA GTTGTCCCTT
198 A C E V T H Q G L S S P V T K S F N R G

721 GAGTGTTAAG CTGATCCTCT ACGCCGGACG CATCGTGGCC CTAGTACGCA ACTAGTCGTA
CTCACAATTC GACTAGGAGA TGCGGCCTGC GTAGCACCGG GATCATGCGT TGATCAGCAT
218 E C O (SEQ ID NO.62)

(SEQ ID NO.65)

FIG. 45

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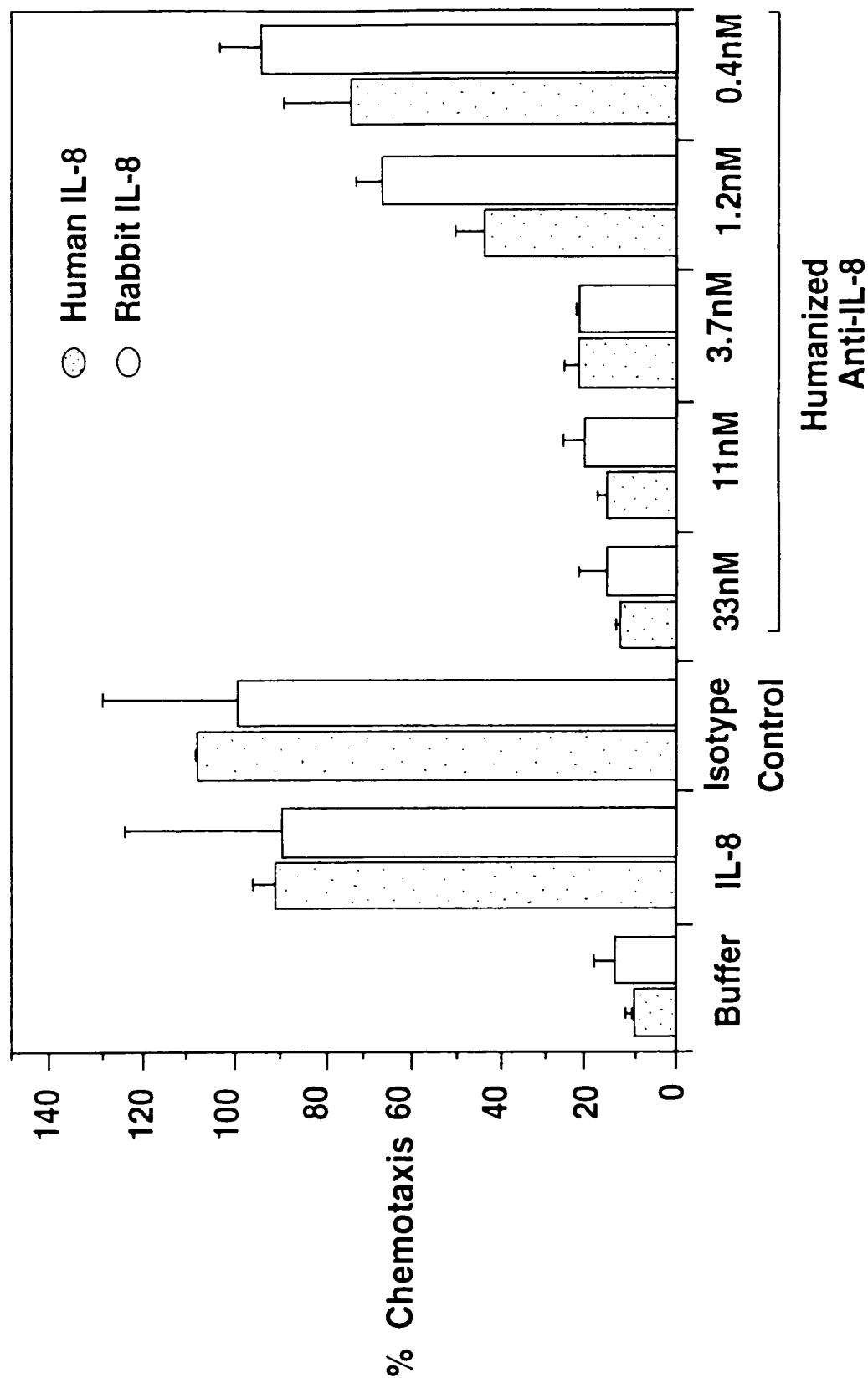


FIG. 46

N35AH1upr
5'-CTAGTGCAGTCTGGCGGTGGCCTGGTGACAGCCAGGGGGCTCACTCCGTTTGTCCTGTGCAGCTTCTGGCTACTCCTTC-3'
(SEQ ID NO.66)

N35AH1lwr
5'-TCGAGAGGAGTAGCCAGAGCTGCACAGGACAAACGGAGTGAGCCCCCTGGCTGCACACAGGCCACCGCCAGACTGCACCT
AG-3' (SEQ ID NO.67)

Bold indicates nucleotide change destroying PvuII site.

FIG. 47

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> length: 8120 (circular)
 >This has the pSVI backbone with the PRK7 cloning linker (pSVI7) and the intron DHFR(ID)
 >made from pSVI.WTSD.D by adding a linearization linker(LL) into the HpaI site

```

cac8I
aluI
sstI
sacI
hgiJII
hgiAI/aspHI
ec1136II
bsp1286
bs1HKAI
bmyI
banII
taqI
1 TTCGAGCTCG CCCGACATTG ATTATTGACT AGAGTCGATC GACAGCTGTG GAATGTGTGT CAGTTAGGGT GTGGAAGTC CCCAGGCTCC CCAGCAGGCA
AAGCTCGAGC GGGCTGTAAC TAATAACTGA TCTCAGCTAG CTCGCGACAC CTTACACACA GTCAATCCCA CACCTTTTCAG GGTCCGAGG GGTGTCGGT

          sau3AI aluI
          mboI/ndeII[dam-]
          dpaI[dam+]
          pvuI/bspCI
          pleI dpaII[dam-]
          hinfI taqI[dam-]
          rmaI mcrI pvuII
          maeI bseI nspBII
          bfaI taqI[dam-]
          bsmFI nlaIV cac8I
          sfaNI
          ppulOI
          nsiI/avaIII
          nlaIII
          sphi
          nspI
          nspHI
          cac8I
          101 GAAGTATGCA AAGCATGCAT CTCAATTAGT CAGCAACCGAG GTGTGGAAG TCCCGAGGCT CCCGAGCAGG CAGAAGTATG CAAAGCATGC ATCTCAATTA
          CTTCATACGT TTCGTACGTA GAGTTAATCA GTCGTTGGTC CACACCTTTC AGGGTCCGA GGGGTCTGCC GTTTCATAC GTTTCGTACG TAGAGTTAAT

          nlaIII
          styI
          ncoI
          bslI dsaI
          aciI bsaJI
          201 GTCAGCAACC ATAGTCCCGC CCCTAACTCC GCCCATCCCG CCCTAACTC CGCCAGTTC CGCCCATCTT CGGCCCATG GCTGACTAAT TTTTTTTATT
          CAGTCGTTGG TATCAGGGCG GGGATTGAG GGGGTAGGCG GGGATTGAG GCGGTCAAG GCGGGTAAGA GCGGGGGTAC CCACTGATTA AAAAAATAA

```

FIG. 48A

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301 TATGAGAGG CCGAGGCCGC CTGGCCTCT GAGCTATTCC AGAAGTAGTG AGGAGGCTTT TTGGAGGCC TAGGCTTTTG CAAAAGCTA GCTTATCCGG
 ATAGTCTCC GGTCCGGCG GAGCCGGAGA CTCGATAAGG TCTTCATCAC TCCTCCGAAA AAACCTCCGG ATCCGAAAC GTTTTTCGAT CGAATAGGCC
 401 CCGGGAACGG TGCATTGGAA CGCGGATTCC CCGTGCCAAG AGTGACGTAA GTACCGCCTA TAGAGCGATA AGAGGATTTT ATCCCGCTG CCATCATGTT
 GGCCCTTGCC ACGTAACCTT CGGCCTAAGG GGCACGGTTC TCACTGCATT CATGGCGGAT ATCTCGCTAT TCCTCTAAA TAGGGCGCAC GGTAGTACCA
 DHER ATG^

FIG. 48B

scrFI
 mvaI
 ecorII
 dsav
 bstNI
 apyI[dcm+]
 sexAI
 tflI
 hinfI
 3del mbolI taqI
 ahalII/draI
 tru9I
 mseI
 ahalII/draI
 601 CAAAGAAATGA CCACAACTC TTCACTGGAA GGTAACAGA ATCTGGTGAT TATGGGTAGG AAAACCTGGT TCTCCATTCC TGAGAAGAAT CGACCTTTAA
 GTTCTTACT GGTCTGGAG AAGTCACCTT CCATTTGTCT TAGACCACTA ATACCCATCC TTTTGGACCA AGAGGTAAGG ACTCTTCTTA GCTGGAATTT
 sstI
 sacI
 hgiIII
 hgiAI/asplI
 eelI36II
 bspI286
 bsiHKA
 bmyI
 mnII aluI
 bssSI banII
 bstXI
 foki
 sfanI mseI
 701 AGGACAGAAT TAATATAGTT CTCAGTAGAG AACTCAAGA ACCACCAAGA GGAGCTCATT TTCTTGCCAA AAGTTTGAT GATGCCCTTA GACTTATTGA
 TCCTGTCTTA ATTATATCAA GAGTCATCTC TTGAGTTTCT TGGTGGTGCT CCTCGAGTAA AAGAACGGTT TTCAACCTTA CTACGGAATT CTGAATAACT
 tru9I
 mseI
 aseI/asnI/vspI
 haeIII/palI
 haeI
 scrFI
 mvaI
 ecorII
 dsav
 bstNI
 nlaIII
 apyI[dcm+]
 hinfI
 apyI[dcm+]
 hinfI
 801 ACAACCGGAA TTGGCAAGTA AAGTAGACAT GGTGGGATA GTCTGTGTTA CCAGGAAGCC ATGAATCAAC CAGGCCACCT TAGACTCTTT
 TGTTGGCCTT AACCGTTCAT TTCATCTGTA CCAAACCTAT CAGCCTCCGT CAAGACAAAT GGTCCTTCGG TACTTAGTTG GTCCGGTGA ATCTGAGAA

FIG. 48C

JAN 22 2003

hgai
 hinli/acyl
 ahali/bsahi
 scrFI
 mval mnlI
 ecorII
 dsav
 bstNI econI
 apyl(dcm+) mnlI
 bsajI bsli ddel
 901 GTGACAAGGA TCATGCAGGA ATTTGAAAGT GACACGTTTT TCCAGAAAT TGATTTGGG AAATATAAAC CTCCTCCAGA ATACCCAGGC GTCCTCTCTG
 CACTGTTCT AGTAGCTCT TAACTTTCA CTGTGCAAAA AGGTCTTTA ACTAAACCC TTTATATTG GAGAGGTCT TATGGGTCCG CAGGAGAGAC

scrFI
 mval
 ecorII
 dsav
 bstNI
 apyl(dcm+)
 sau96I
 avall

sfanI mnlI mnlI alul
 1001 AGTCCAGGA GGAAGAAGC ATCAAGTATA AGTTTGAAGT CTACGAGAAG AAAGACTAAC AGGAAGATGC TTCAAGTTC TCTGTCCCC TCCTAAAGT
 TCCAGGTCT CCITTTTCCG TAGTTCATAT TCAAACTTCA GATGCTCTTC TTTCTGATTG TCCTCTCAGG AAAGTTCAAG AGACGAGGG AGGATTTTCA
 ^END DHFR

styl
 bsajI
 sau3AI
 mboI/ndeII(dam-)
 dpnI(dam+)
 dpnII(dam-)
 alwI(dam-)
 bstYI/xhoII
 cac8I
 ppulOI
 nsII/avallI
 1101 ATGCATTTT ATAGACCAT GGGACTTTTG CTGGCTTTAG ATCCCTTGG CTTCGTTAGA ACGCAGCTAC AATTAATACA TAACCTTATG TATCATACAC
 TACGTAAAA TATTCTGGTA CCTGTGAAAC GACCGAAATC TAGGGGAACC GAAGCAATCT TCGGTGATG TTAATTATGT ATTGGAATAC ATAGTATGTG


sau96I
 avall
 asul
 scrFI
 mval
 ecorII

FIG. 48D

1

FIG. 48E

FIG. 48F



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scrFI
 mvaI
 ecorII
 ecoNI
 dsav
 bstNI
 bslI
 apyI[dcn+]
 fnu4HI
 bsoFI
 bsvI
 bslI
 agel
 tthlII/aspI
 ddel
 hhaI/cfoI
 nspBII
 alw44I/snoI
 cauII
 scfI
 1801 CTGGGCTGCC TGGTCAAGGA CTACTCCCG GAACCGGTGA CGGTGCTGTG GAACTCAGG GCCCTGACCA CGGGGTGCA CACCTTCCCG GGTGTCTTAC
 GACCCGACGG ACCAGTTCTT GATGAGGGG CTTGGCCTT GCCACAGAC CTTGAGTCCG CGGACTGGT CGCCGACCT GTGGAAGGCG CGACAGGATG
 147 L G C L V K D Y F P E P V T V S W N S G A L T S G V H T F P A V L Q
 fnu4HI
 bsoFI
 rmal
 bsp1286
 maeIII
 hphI
 bmyI
 mnlI
 bbvI
 ddeI
 fnu4HI
 bsoFI
 mnlI
 hinfI
 ddeI
 pleI
 mnlI
 hinfI
 eco8II
 bsu36I/mstII/sauI
 ddeI
 CTCAGCAGG
 TGGTACTCT
 ACTTACTCC
 TGAGATGAGG
 GAGTCGTCC
 ACCACTGACA
 CGGGAGATCG
 TCGAACCCGT
 GGGTCTGGAT
 GTAGACCTTG
 CACTTACTGT
 TCGGGTCTT
 181 S S G L Y S L S S V V T V P S S S L G T Q T Y I C N V N H K P S N
 hgiJII
 bsp1286
 bmyI
 banII
 maeIII
 nspI
 nspHI
 nlaIII
 scrFI
 mvaI
 ecorII
 dsav
 bstNI
 bsaJI
 mslI
 2001 CACCAAGGTG GACAGAAG TTAGGCCCCA ATCTTGTGAC AAAACTGACA CACTGCCACC GTGCCAGCA CCTGAACCTC TGGGGGACC GTCAGTCTTC
 GTGGTCCAC CTGTCTTTT AACTCTTTT GAGTCTTCTT TTTTGTGTGT TAGAACAATG TTTTGTGTGT CTACGGGTGG CACGGGTCTG GGACTTGAGG ACCCCCTGG CACTCAGAAC
 214 T K V D K K V E P K S C D K T H T C P C P A P E L L G G G P S V F
 ahdI/eam1105I
 sau96I
 avaII
 scrFI
 mvaI
 ecorII
 dsav
 bstNI
 bsaJI
 mslI
 2001 CACCAAGGTG GACAGAAG TTAGGCCCCA ATCTTGTGAC AAAACTGACA CACTGCCACC GTGCCAGCA CCTGAACCTC TGGGGGACC GTCAGTCTTC
 GTGGTCCAC CTGTCTTTT AACTCTTTT GAGTCTTCTT TTTTGTGTGT TAGAACAATG TTTTGTGTGT CTACGGGTGG CACGGGTCTG GGACTTGAGG ACCCCCTGG CACTCAGAAC
 214 T K V D K K V E P K S C D K T H T C P C P A P E L L G G G P S V F

FIG. 48G



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```
sau96I
nlaIV
mspI
hpaII
scrFI
ncII
dsav
sau3AI auaII nlaIII
mboI/ndeII[dam-] nspI
nlaIII cauII mnII nspHI
rcaI dpaI[dam+] ddel mslI
mnII dpaII[dam-] eco8II maeIII
styI mslI bsaJI asuI bsu36I/mstII/sauI
earI/ksp632I bsaJI mslI bspHI[dam-] asuI bsu36I/mstII/sauI
2101 CTCTTCCCC CAAAACCCAA GGACACCCCTC ATGATCTCCC GGACCCCTGA GGTACATGC GTGGTGGTGG ACCTGAGCCA CGAAGACCCT GAGGTCAAGT
GAGAAGGGGG GTTTGGGT CCGTGGGG TACTAGAGG CCGGGGACT CCACTGTACG CACCACACC TGCACCTGGT GCTTCTGGGA CTCCAGTTCA
247 L F P P K P K D T L M I S R T P E V T C V V V D V S H E D P E V K F
drdI mnII
mboII ddel
bpuAI eco8II
bbsI bsu36I/mstII/sauI
maeII
rsal
csp6I
hphI
hgaI mnII
GTGGTACCGT GTGGTACCGT GTGGTACCGT GTGGTACCGT GTGGTACCGT GTGGTACCGT GTGGTACCGT GTGGTACCGT GTGGTACCGT GTGGTACCGT
2201 TCAACTGGTA CGTGGACGGC ATGATGCCAA GACAAAGCCG CGGGAGGAGC AGTACACAG CAGTACCGT CAGTACCGT CAGTACCGT CAGTACCGT CAGTACCGT
AGTTGACCAT GCACCTGCCG CACCTCCACG TATTACGGT CTGTTGGC GCCCTCTCG TCATGTTGTC GTGATGGCA CACCAGTCGC AGGAGTGGGA
281 N W Y V D G V E V H N A K T K P R E E Q Y N S T Y R V V S V L T V
maeII
rsal
csp6I
bsrI bsaI
mnII
bsaI
rsal
csp6I
hphI
hgaI mnII
GTGGTACCGT GTGGTACCGT GTGGTACCGT GTGGTACCGT GTGGTACCGT GTGGTACCGT GTGGTACCGT GTGGTACCGT GTGGTACCGT GTGGTACCGT
2201 TCAACTGGTA CGTGGACGGC ATGATGCCAA GACAAAGCCG CGGGAGGAGC AGTACACAG CAGTACCGT CAGTACCGT CAGTACCGT CAGTACCGT CAGTACCGT
AGTTGACCAT GCACCTGCCG CACCTCCACG TATTACGGT CTGTTGGC GCCCTCTCG TCATGTTGTC GTGATGGCA CACCAGTCGC AGGAGTGGGA
281 N W Y V D G V E V H N A K T K P R E E Q Y N S T Y R V V S V L T V
scrFI
mvaI
ecoRII
dsav
ecoNI bstNI bsrI
bsI apyI[dcmt+]
2301 CCTGCACCCAG GACTGGCTGA ATGGCAGGA GTACAAGTGC AAGTCTCCA ACAAGCCCT CCCAGCCCCC ATCGAGAAA CCATCTCAA AGCCAAAGGG
GGACGTGGTC CTGACCGACT TACCGTCTCT CATGTTACG TTCCAGAGGT TGTTCGGGA GGTCCGGGG TAGTCTTTT GGTAGAGTT TCGGTTTCCC
314 L H Q D W L N G K E Y K C K V S N K A L P A P I E K T I S K A K G
fnu4HI
bsoFI
bbvI
```

FIG. 48H

FIG. 48I

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```

sau96I      acII      haeIII/palI
          fnu4HI      asuI
          bsoFI      nlaIII
          sfiI      styI      aluI
          eaeI      ncoI      fnu4HI
          cfrI      dsal      bsoFI
          aluI      haeIII/palI      bbvI      maeIII
          asuI      bfaI      accI      bspMI      hindIII      bgli      bsaJI
          bsmAI      cauII
2701 TCCGTGTC CGGTAATG AGTGGAGCGG CCTAGAGTC GACCTGCAGA AGCTGGCCG CCATGGCCCA ACTTGTTTAT TGCAGCTTAT AATGGTTACA
AGGACAGAG GCCAATTAC TCACGCTGCC GGGATCTCAG CTGGACGTCT TCGAACCGGC GGTACCGGGT TGAACAATA ACGTGGAATA TTACCAATGT
447 S L S P G K O (SEQ ID NO.71)

          rmaI
          maeI
          bsmI      bfaI
          sfanI      apoI
2801 AATAAGCAA TAGCATCACA AATTTCACAA ATAAAGCAAT TTTTCACTG CATCTAGTT GTGGTTTGTG CAAACTCATC AATGTATCTT ATCATGTCTG
TTATTTCGTT ATCGTAGTGT TTAAGTGTT TATTTCGTAA AAAAAGTGAC GTAAGATCAA CACCAACAG GTTTGAGTAG TTACATAGAA TAGTACAGAC
          nlaIII      alwI(dam-)

sau3AI      mboI/ndeII(dam-)
          dpnI(dam+)
          dpnII(dam-)
          pvuI/bspCI
          mcrI
          bseI
          taqI(dam-)      tru9I
          claI/bspl06(dam-)
          bspDI(dam-)      mseI
          sau3AI      xmnI
          mboI/ndeII(dam-)
          dpnI(dam+)      asp700
          dpnII(dam-)      asel/asnI/vspI      bsaJI
          hhaI/cfoI      nlaIII
          hinPI      dsal      haeIII/palI
          fnu4HI      haeI
          bsoFI      styI
          bbvI      ncoI
          mnlI      mnII
          asp718      mnlI
          acc65I      ddeI      acII
2901 GATCATCGG GAATTAATTC GCGGAGCAGC CATGGCCTGA AATAACCTCT GAAAGAGGAA CTGGTTAGG TACCTTCTGA GCGGGAAGA ACCATCTGTG
CTAGTAGCC CTTAATTAG CCGCGTCGTG GTACCGGACT TTATTGGAGA CTTTCTCCTT GAACCAATCC ATGGAAGACT CCGCCTTCTT TGTAGACAC

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FIG. 48J

FIG. 48K

FIG. 48L

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[illegible]

FIG. 48M

FIG. 48N

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FIG. 480

FIG 48P

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```

nlaIII
styl
ncol
bsli dsal
acil bsajI
CGCCCCATG GGTGACTAAT
TAGAGTTAAT CAGTCGTTGG TATCAGGGCG GGGATTGAG GGGGTAGAG GGGGGGTAC CGACTGATTA

acil
bsmFI
acil fokI
acil bsrI acil
CGCCTAACTC CGCCAGTTC CGCCCATCTT CGCCCATCTT GGTGACTAAT
TAGAGTTAAT CAGTCGTTGG TATCAGGGCG GGGATTGAG GGGGTAGAG GGGGGGTAC CGACTGATTA

fnu4HI
bsoFI
bglI
sfII
haeIII/palI
mnII mnII ddeI
haeIII/palI bsajI mnII aluI
mnII bsajI acil haeIII/palI
CGGAGGCGG CTGGCCTCT GAGCTATTCC AGAAGTAGTG AGGAGGCTTT TTTGGAGGCC ATCGGAAAC GTTTTTCGAC
AAAAAATAA ATACGTCTCC GGCTCGGGCG GAGCGGAGA CTGGATAAGG TCTTCATCAC TCCTCGGAAA AAACCTCCGG ATCGGAAAC GTTTTTCGAC
start pUC118^

fnu4HI
haeIII/palI hinPI
mcrI hhaI/cfoI
eagI/xmaIII/eclXI thal
eaeI fnuDII/mvnI
notI
bsrBI bsoFI hinPI bspMI
taqI cfrI hhaI/cfoI scfI
xhoI fnu4HI tru9I cac8I tru9I pstI
paeR7I bslEI pacI ascI ahaIII/draI
avaI bsoFI msel tru9I bshl236I msel bsgI maeIII cfrI
mnII acil acil maeI bssHII swaI sse8387I aluI bsrI
CGGCGGCTTA ATTAAGGCGG GCCATTAA TCCTGAGGT AACAGCTGG CACTGGCGCT CGTTTACAA CGTCGTGACT GGGAAACCC
AATGGAGCTC GCCGGCAAT TAATTCGGG CGGTAAATTT AGGAGCTCCA TTGTGGAACC GTGACCGGCA GCAAAATGTT GCAGCACTGA CCCTTTTGGG
^linearization linker inserted into HpaI site

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FIG. 48Q

FIG. 48B

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```

mspI      nlaIV      hgiCI      taqI
hpaII     hgiJII     baniI      mnlI
nael      bsp1286     bmyI
cfr101/bsrFI      aluI      nlaIV
maeII     cac8I      dralII     sau96I      hphI      maeII     pleI      tru9I      pleI
                    bsaAI     asuI                    drdI     hinfi     maeII     msel     hinfi
5301 TTTCTCGGCA CGTTGCGCGG CTTTCCCGCT CAAGCTCTAA ATCGGGGGCT CCCTTTAGG TTCCGATTTA GTGCTTTACG GCACCTCGAC CCCAAAAAAC
AAAGAGCGGT GCAAGCGGCC GAAAGGGCA GTTCGAGATT TAGCCCCCGA GGGAAATCCC AAGGTAAAT CAGGAATGC CGTGAGCTG GGGTTTTTTG
                    maeII     haeIII/palI
                    dralII     sau96I
                    bsaAI     asuI
hphI      maeII     pleI      tru9I      pleI
5401 TTGATTTGGG TGATGGTTCA CGTAGTGGG CATCGCCCTG ATACACGGTT TTTCGCCCTT TGACGTTGGA GTCCACGTTT TTTAATAGTG GACTCTTGT
AACTAAACCC ACTACCAAGT GCATCACCCG GTAGCGGAC TATCTGCCAA AAAGCGGAA ACTGCAACCT CAGGTGCAAG AAATTATCAC CTGAGAACAA
                    bslI      bslI     auaI      haeIII/palI      aluI     msel     tru9I     msel
bsrI      bslI     auaI      haeIII/palI      aluI     msel     tru9I     msel
5501 CCAAACTGGA ACAACTCA ACCCTATCTC GGGCTATTCT TTTGATTTAT AAGGGATTTT GCGGATTTGG GCCTATTGGT TAAAAATGA GCTGATTTAA
GGTTTGACCT TGTTCGAGT TGGGATAGAG CCGGATAAGA AACTAAATA TTCCCTAAAA CGGCTAAGC CGGATAACCA ATTTTACT CGACTAAAT
                    thal      fnuDII/mvnl      maeII     psp1406I      tru9I     msel     apoI     bsh1236I      sspl     msel
                    tru9I     apoI     tru9I      msel     bsh1236I      sspl     msel
5601 CAAAAATTTA ACGCGAATTT TAACAAATA TTAACGTTTA CAATTTTATG GTGCACCTCTC AGTACAATCT GCTCTGATGC CGCATAGTTA AGCCAACCTCC
GTTTTTAAAT TGGGCTTAA ATTTGTTTAT AATTGCAAT GTTAAAAATAC CACGTGAGAG TCATGTTTGA CGAGACTACG GCGTATCAAT TCGGTTGAGG
                    hinPI      hhaI/cfoI      thal      fnuDII/mvnl      bstUI      nspBII     bsh1236I      aciI     hgal     drdI
                    maeII     bsrI     nlaIII     hhaI/cfoI      fnu4HI     bsoFI      maeII     bsh1236I      sspl     msel     apoI     bsh1236I      sspl     msel
                    bsaAI     thlIII/aspI     bbvI      bsaAI     thlIII/aspI     bbvI
5701 GCTATCGCTA CGTGACTGGG TCATGGCTGC GCGCCGACAC CCGCTGACGC GCGCTGACGC GCTTGCTGTC TCCCGGCATC CGCTTACAGA
CGATAGCGAT GCACTGACCC ACTACCGACG CCGGCTGTG GCGGCTGTG GCGGCTGTG GCGGCTGTG CGAACAGACG AGGCGCGTAG CGGAATGTCT

```

FIG. 48S

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FIG. 48T

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```

6201 bssSI maeIII taqI alwI[dam-] aciI bstYI/xhoII
      CACGAGTGGG TTACATCGAA CTGGATCTCA ACAGCGGTAA GATCCTTGAG AGTTTCGGC CCGAAGAAGC TTTTCCAATG ATGAGCACTT TTAAGTTCT
      GTGCTCACC AATGTAGCTT GACCTAGAGT TGTCGCCATT CTAGGAAGCTC TCAAGAGCGG GCGTCTTTCG AAAGGTTAC TACTCGTGAA AATTCAAGA

      sau3AI nspBII sau3AI mboI/ndeII[dam-] maeII
      mboI/ndeII[dam-] dpnI[dam+] psp1406I
      dpnI[dam+] dpnII[dam-] xmnI
      bstYI/xhoII alwI[dam-] asp700
      bsrI dpnII[dam-] alwI[dam-] mboII

      scrFI
      aciI
      thaI
      fnuDII/mvnI
      bstUI
      bshI236I
      hinPI
      hhAI/cfoI
      GGTATGTGC GCGGTATTAT CCGTGTATGA CGCGGGGCAA GAGCAACTCG GTCGCGGCAT ACACATTCTT CAGATGACT TGGTTGAGTA CTCACCACTC
      GGATACACCG CGCCATAATA GGGCACTACT GCGGCCCGTT CTCGTTGAGC CAGCGGGCGTA TGTGATAAGA GTCTTACTGA ACCAACTCAT GAGTGGTCAG

      aciI
      nciI
      mspI
      hpaII
      dsav
      hinII/acyI
      hgaI caulI
      ahaII/bseHI
      bcoI bsiFI bsoFI ddel
      mcrI fnu4HI
      bsiFI
      rsal
      csp6I bsrI
      scaI hphI maeIII
      TGGTTGAGTA CTCACCACTC
      TGTGATAAGA GTCTTACTGA ACCAACTCAT GAGTGGTCAG

      sau3AI
      mboI/ndeII[dam-]
      dpnI[dam+]
      dpnII[dam-]
      pvul/bspCI
      mcrI
      bsiFI
      haeIII/palI
      eaeI
      cfrI
      fnu4HI
      bsoFI
      aciI
      haeIII/palI
      eaeI
      cfrI
      fnu4HI
      bsoFI
      aciI
      sau3AI maeIII mspI
      mboI/ndeII[dam-] sau3AI nlaIV
      dpnI[dam+] mboI/ndeII[dam-] aluI
      dpnII[dam-] dpnI[dam+] hpaII
      nlaIII
      sau3AI maeIII mspI
      mboI/ndeII[dam-]
      dpnI[dam+]
      dpnII[dam-]
      nlaIII alwI[dam-] bsaWI
      aluI aciI
      ACCGTTTTT TGCACACAT GGGGGATCAT GTAACCTGCGC TTGATCGTTG GGAACCGGAG CTGAATGAAG CCATACCAAA
      ACCCTCCTGG CTTCCTCGAT TGGCGAAAAA ACGTGTGTGA CCCCCTAGTA CATTGAGCGG AACTAGCAAC CCTTGGCCTC GACTTACTTC GGTATGGTTT

      sau96I
      auaII
      asuI
      mnlI
      aluI
      aciI
      TCGGAGGACC GAAGGAGCTA ACCGTTTTT TGCACACAT GGGGGATCAT GTAACCTGCGC TTGATCGTTG GGAACCGGAG CTGAATGAAG CCATACCAAA
      ACCCTCCTGG CTTCCTCGAT TGGCGAAAAA ACGTGTGTGA CCCCCTAGTA CATTGAGCGG AACTAGCAAC CCTTGGCCTC GACTTACTTC GGTATGGTTT

```

FIG. 48U

hinPI mspI
 hhai/cfoI hpaII
 mstI aluI scrFI
 avilI/fspI bsrI tru9I
 maeII maeI dsav msel
 psp1406I cauII asel/asnI/vspI
 6601 CGACGAGCGT GACACCACGA TGCCAGCAGC AATGGCAACA AACTATTAACT TGGCGAACTA CTTACTCTAG CTTCCCGGCA ACAATTAATA
 GCTGCTCGCA CTGTGTGTCT ACGGTGCTCG TTACCGTTCT TGAACGCGT TGGATAATTG ACCGCTTGAT GAATGAGATC GAAGGCCCGT TGTTAATTAT
 bglI mspI
 sau96I cac8I
 haeIII/palI hpaII
 hinPI asuI mspI
 hhai/cfoI hpaII
 6701 GACTGGATGG AGGCGGATAA AGTTGAGGA CCACTTCTGC GCTCGGCCCT TCCGGCTGGC TGGTTTATTG CTGATAAATC TGGAGCCGGT GAGCGTGGGT
 CTGACCTACC TCCGCTATT TCAACGTCCT GGTGAAGACG CGAGCCGGGA AGCCGACCG ACCAAATAAC GACTATTTAG ACCTCGGCCA CTGCGACCCA
 aciI
 fnu4HI haeIII/palI
 fnuDII/mvnI bsoFI sau96I
 bstUI nlaIV
 bsh1236I bsrDI bsrI asuI mnlI
 6801 CTCGGGGTAT CATTCGACGA CTGGGGCCAG ATGGTAAGCC CTCGCTATC GTAGTTATCT ACACGACGGG GACTCAGGCA ACTATGGATG AACGAAATAG
 GAGGCCCATTA GTAACGTCGT GACCCCGGTC TACCATTCCG GAGGCGATAG CATCAATAGA TGTGCTGCC CTGAGTCCGT TGATACCTAC TTGCTTTATC
 ddeI
 sau3AI nlaIV
 mboI/ndeII[dam-]
 dpnI[dam+] hgICI tru9I msel tru9I
 dpnII[dam-] bani mnlI maeII
 6901 ACAGATCGCT GAGATAGGTG CCTCACTGAT TAAGCATTTGG TAACTGTGAG ACCAAGTTTA CTCATATATA CTTTAGATTG ATTTAAAACT TCATTTTAA
 TGTTAGCGA CTCTATCCAC GGAGTGACTA ATTGCTAACC ATTGACAGTC TGGTTCAAAT GAGTATATAT GAAATCTAAC TAAATTTTGA AGTAAAAAT
 rnaI sau3AI
 maeI mboI/ndeII[dam-]
 sau3AI hpaI mboI/ndeII
 7001 TTTAAAGGA TCTAGTGAA GATCCTTTT GATAATCTCA TGACCAAAAT CCCTTAACGT GAGTTTTCGT TCCACTGAGC GTCAGACCCC GTAGAAAAGA
 AAATTTTCCT AGATCCACTT CTAGGAAAA CTATTAGAGT ACTGGTTTTA GGGAAATTGCA CTCAAAAGCA AGGTGACTCG CAGTCTGGGG CATCTTTCT

FIG. 48V



```

sau3AI
mboII[dam-]
sau3AI mboI/ndeII[dam-] thaI
mboI/ndeII[dam-] fnuDII/mvni
dplI[dam+] dplI[dam+] bstUI cac8I
dplII[dam-] dplII[dam-] bsh1236I fnu4HI
bstYI/xhoII alwI[dam-] hinPI bsoFI
alwI[dam-] bstYI/xhoII hhal/cfoI bbvI
7101 TCAAAGGATC TTCTTGAGAT CCTTTTTC TGCGGGTAAT CTGCTGCTTG CAAACAAAAA AACACCGCT ACCAGCGGTG GTTGTGTTGC CGGATCAAGA
AGTTTCCTAG AAGAAGCTCTA GGAAGAAAAAG ACGCGCATTA GACGACGAAC GTTGTGTTT TGTGTGCGCA TGGTCGCCAC CAAACAAACG GCCTAGTTCT
          acII nspBII hpaII aluI
          mspI
          rmaI maeI haeIII/palI
          bfaI bslI haeI
7201 GCTACCAACT CTTTTTCGA AGGTAAGCTGG CTTACAGCAGA GCGCAGATAC CAAATACTGT CCTTCTAGTG TAGCCGTAGT TAGGCCACCA CTTCAAGAAC
CGATGGTTGA GAAAAGGCT TCCATTGACC GAAGTGTCT CCGCTCTATG GTTATGACA GGAAGATCAC ATCGGCATCA ATCGGTGCTG GAAGTTCTTG
          fnu4HI
          bsoFI
          bbvI
          fnu4HI
          alwNI[dam-]
          bsrI bsoFI
          maelII bbvI bsrI
          scfI acII mnlI
          bsrI bsoFI
          maelII bbvI bsrI
          bsrI bsoFI
          pleI
          hinfI
7301 TCTGTAGCAC CGCTACATA CCTCGCTCTG CTAATCCTGT TACCAGTGGC TGCTGCCAGT GCGGATAAGT CGTGTCTTAC CGGTTGGAC TCAAGACGAT
AGACATCGTG GCGGATGTAT GGAGCGAGAC GATTAGACA ATGGTCACCG ACGACGGTCA CCGCTATTCA GCACAGAATG GCCCAACCTG AGTTCTGCTA
          hgiAI/asphI
          bsp1286
          bsiHKA
          bmyI
          apaLI/snoI
          alw4I/snoI aluI
          maelII hhal/cfoI
          bsrI bsoFI
          bbvI mcrI
          hinPI bsiEI
          bsaWI
          maelII
          ddeI scfI
7401 AGTTACCGGA TAAGCGCAG CGGTGGGCT GAACGGGGG TCGTGGACA CAGCCAGCT TGGAGCGAAC GACCTACACC GAAGTACGAT ACCTACAGCG
TCAATGGCCT ATTCCGCGTC GCCAGCCCGA CTTGCCCGCC AAGCAGCTGT GTCCGGTCA ACCTCGCTTG CTGGATCTTA TGGATCTGCG

```

FIG. 48W

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7501 TGAGCATTTGA GAAAGCGCA CGCTTCCCGA AGGAGAAAG GCGGACAGGT ATCCGGTAAAG GCGCAGGGTC GGAACAGGAG AGCGCACGAG GGAGCTTCCA
 ACTCGTAACT CTTTCGGGT GCGAAGGGCT TCCCTCTTTC GGCCTGTCCA TAGGCCATTC GCGTCCCG CTTGTCTC TCGGTGCTC CCTCGAAGGT
 scrFI mvaI ecorII dsav bstNI
 hinPI fnu4HI
 hhaI/cfoI bssSI
 haeII bslI bsoFI
 acII bsaWI acII
 hhaI/cfoI aluI apyI(dcm+)

7601 GGGGNAACG CTGGTATCT TTATAGTCCT GTCGGGTTTC GCCACCTCTG ACTGAGCGT CGATTTTGT GATGCTCGTC AGGGGGCGG AGCCTATGGA
 CCCCCCTTGC GGACCATAGA AATATCAGGA CAGCCCAAAG CCGTGGAGAC TGAACCTGCA GCTAAACA CTACGAGCAG TCCCCCGCC TCGGATACCT
 scrFI mvaI ecorII dsav bstNI
 haeIII/palI
 apyI(dcm+) mnlI drdI hgaI
 haeIII/palI haeIII/palI
 scrFI mvaI bslI
 ecorII
 acII
 thal bslI
 fnuDII/mvnl
 bstUI
 cac8I bsh1236I
 nlaIV haeI
 apyI(dcm+) haeI
 haeIII/palI nspI
 haeI
 ac8I
 aflIII
 tfil
 hinfi
 nlaIV

7701 AAAAGCCCGG CAACGGCGG TTTTACGGT TCCTGGCCTT TTGCTGGCT TTTGCTCACA TGTTCTTTCC TCGGTTATCC CTGTATTCTG TGGATAACCG
 TTTTGGGTC GTTGGCGG AAAAATGCCA AGGACCGGAA AACGACCGA AACGAGTGT ACAGAAAGG ACGCAATAGG GGAATAAG ACCTATTGGC
 fnu4HI
 bsoFI
 bbvI
 cac8I acII
 bsrBI fnu4HI
 acII bsoFI
 aluI
 hhaI/cfoI
 mcrI
 bslI
 hhaI/cfoI
 mnlI
 acII
 haeII
 sapI
 hinPI
 mboII hhaI/cfoI
 earI/ksp632I

7801 TATTACCGG TTTGAGTGAG CTGATACCG TCGCCGCGAG CGAAGCAGG GTGAGTGAGC GAGGAGCGG AAGAGCGCC AATACGCAAA
 ATAATGCGG AACTCACTC GACTATGGG AGCGGCGTGC GTTGTGTC TCGGTGCT CAGTCACTCG CTCCTTCGCC TTCTCGCGG TTATGGGTTT
 acII
 aluI
 hhaI/cfoI
 mcrI
 bslI
 hhaI/cfoI
 mnlI
 acII
 haeII
 sapI
 hinPI
 mboII hhaI/cfoI
 earI/ksp632I

FIG. 48X

FIG. 48Y



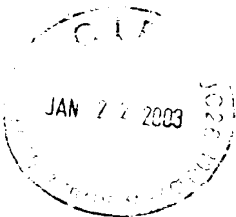
```
tru9I
mseI
aseI/asnI/vspI
xmni
nlaIII asp700
8101 TGACCATGAT TAGGAATTAA (SEQ ID NO.68)
ACTGGTACTA ATGCTTAATT

>length: 8120

aatII(GACGTC): 1690 5947
acc65I(GGTACC): 2969 3967 4529
accI(GTMRAC): 823 1039 2738 4237
acII(CCGC): 217 229 238 250 260 271 317 422 454 485 574 1385 1795 1871 2248 2250 2758 2982
3167 3179 3188 3200 3210 3221 3267 3372 3404 3449 3686 3949 4021 4318 4542 4727
4739 4748 4760 4770 4781 4827 4910 4914 5070 5127 5153 5166 5203 5217 5220 5248
5275 5680 5699 5741 5751 5790 5979 6026 6125 6234 6311 6355 6476 6522 6713 6804
7166 7175 7310 7420 7541 7560 7687 7715 7806 7827 7834 7877 7901 7911 7967 8070

see hinII
acyI
afIII/bfII(CTTAAG): 786
afIII(ACRYGT): 932 7758
ageI(ACCGGT): 1833
ahaII/baII(GRCGYC): 988 1690 1858 5117 5947 6329
ahaII/draI(TTTAAA): 696 4935 6290 6982 7001
ahdI/eamII05I(GACNNNNNGTC): 2087 6865
alul(AGCT): 5 44 332 386 390 753 1097 1165 1370 1431 1951 2603 2751 2784 3282 3336 3340
3562 3566 3676 3733 3792 4270 4288 4311 4344 4554 4842 4896 4954 5047 5333 5590
5803 5822 6516 6579 6679 7200 7457 7593 7819 7937 8096
alw44I/snoI(GTGCAC): 1876 5651 6198 7444
```

FIG. 48Z



APPROVED	O.G. FIG.
BY	CLASS
DRAFTSMAN	SUBCLASS

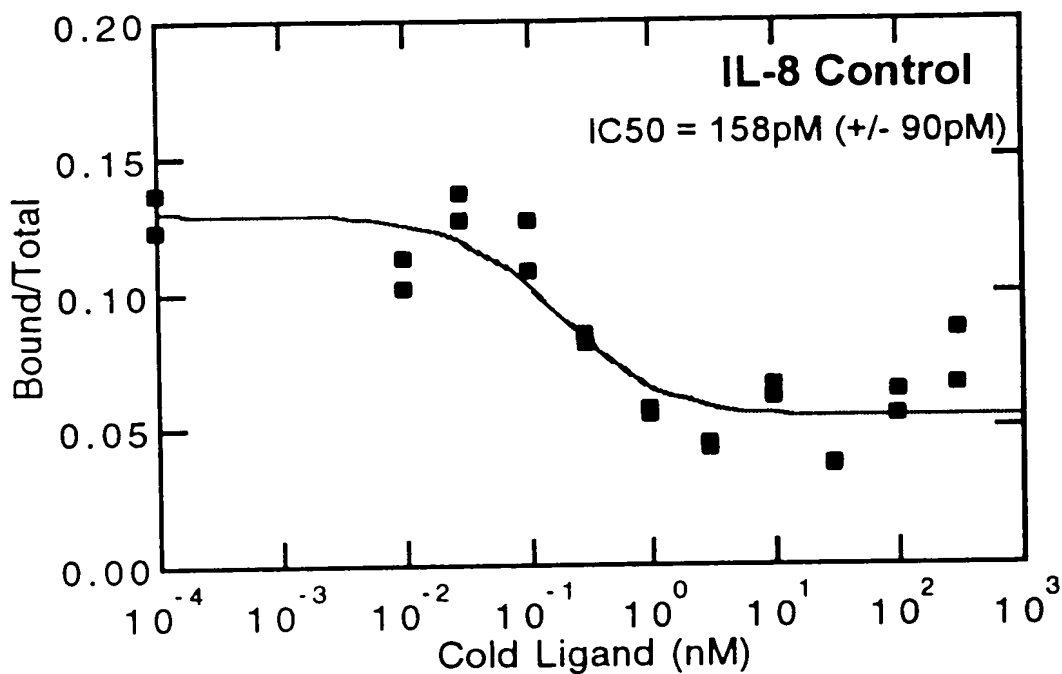


FIG. 49A

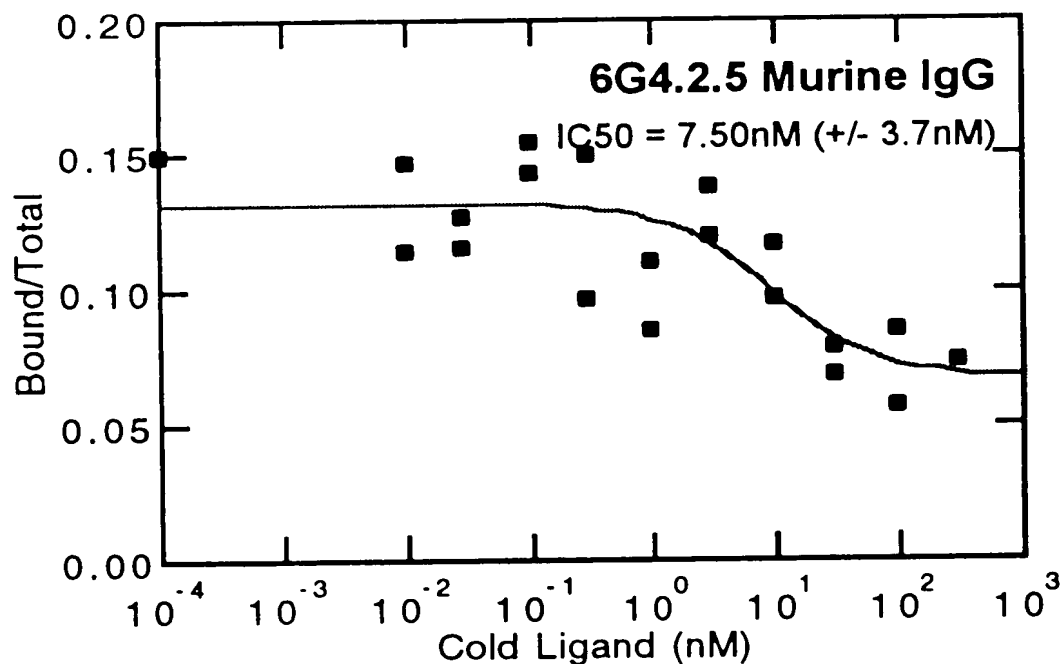


FIG. 49B

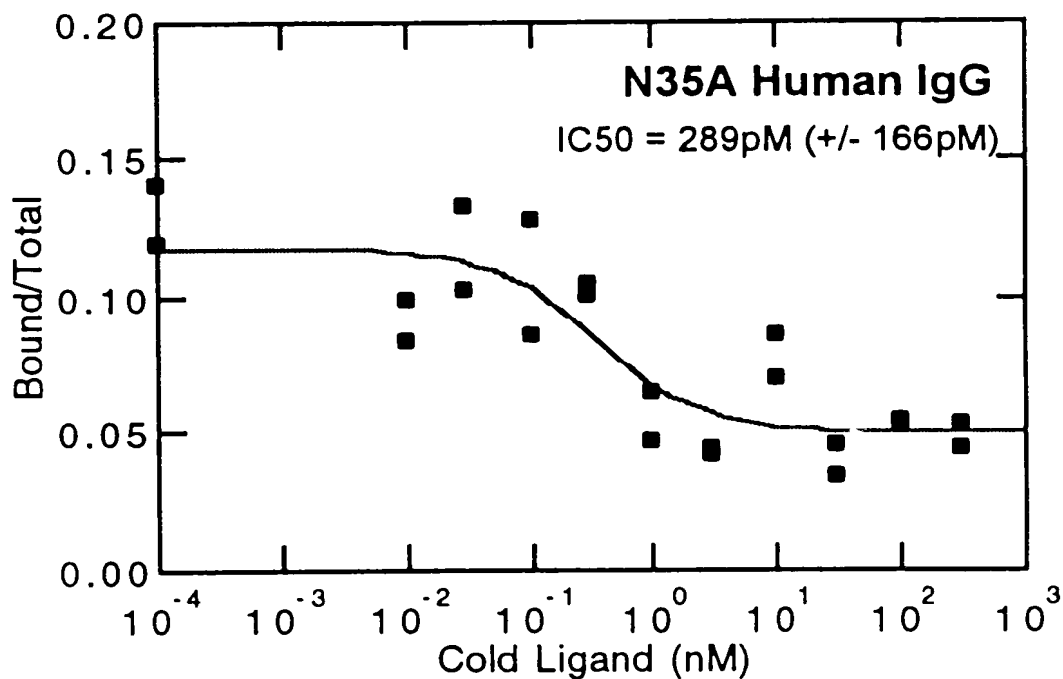
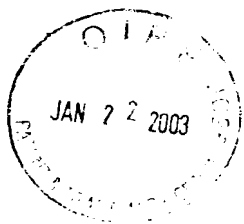


FIG. 49C

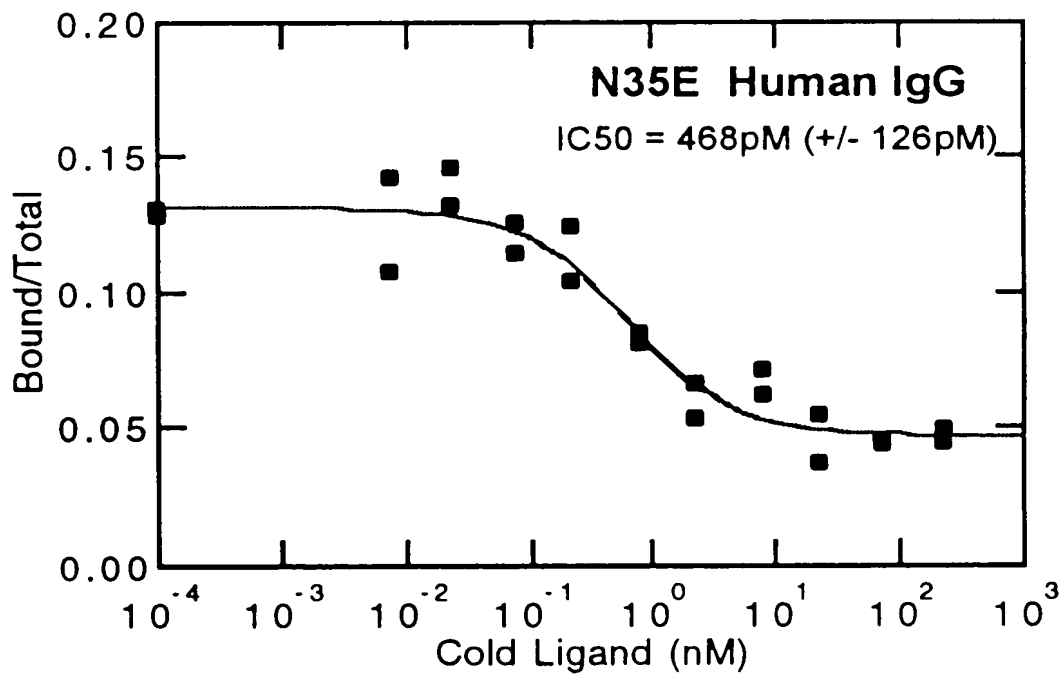


FIG. 49D

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JAN 22 2003

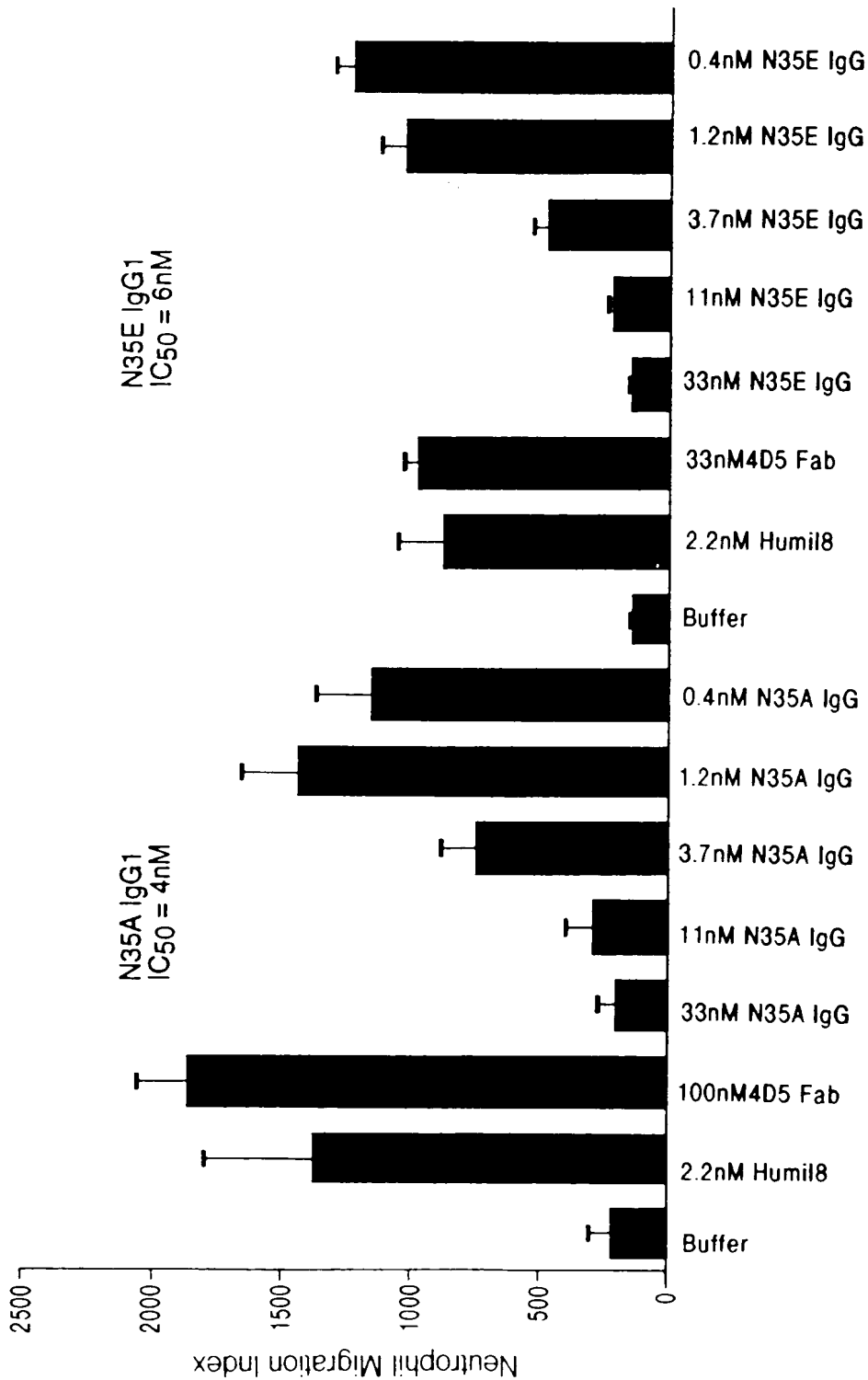


FIG. 50A

JAN 22 2003

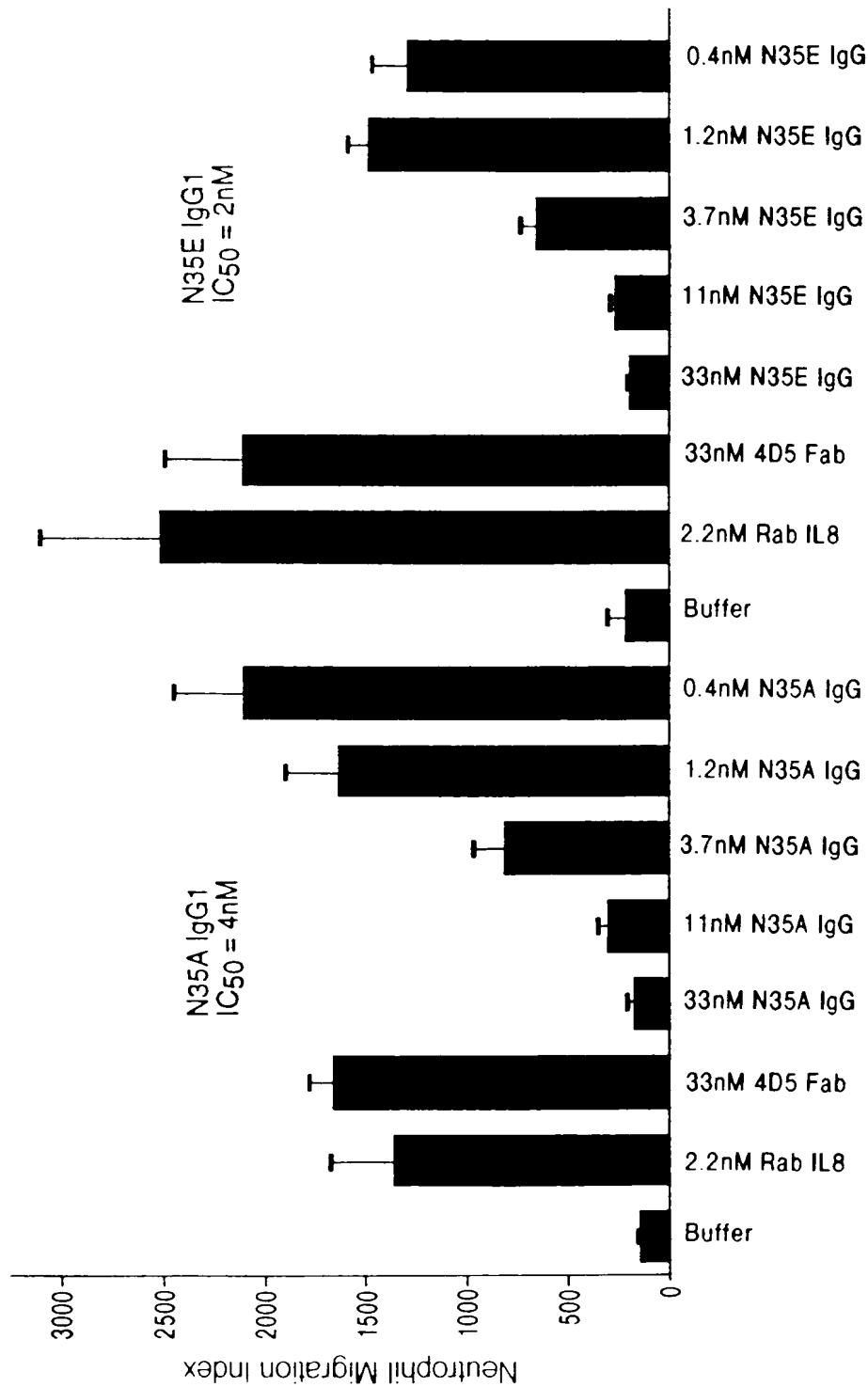
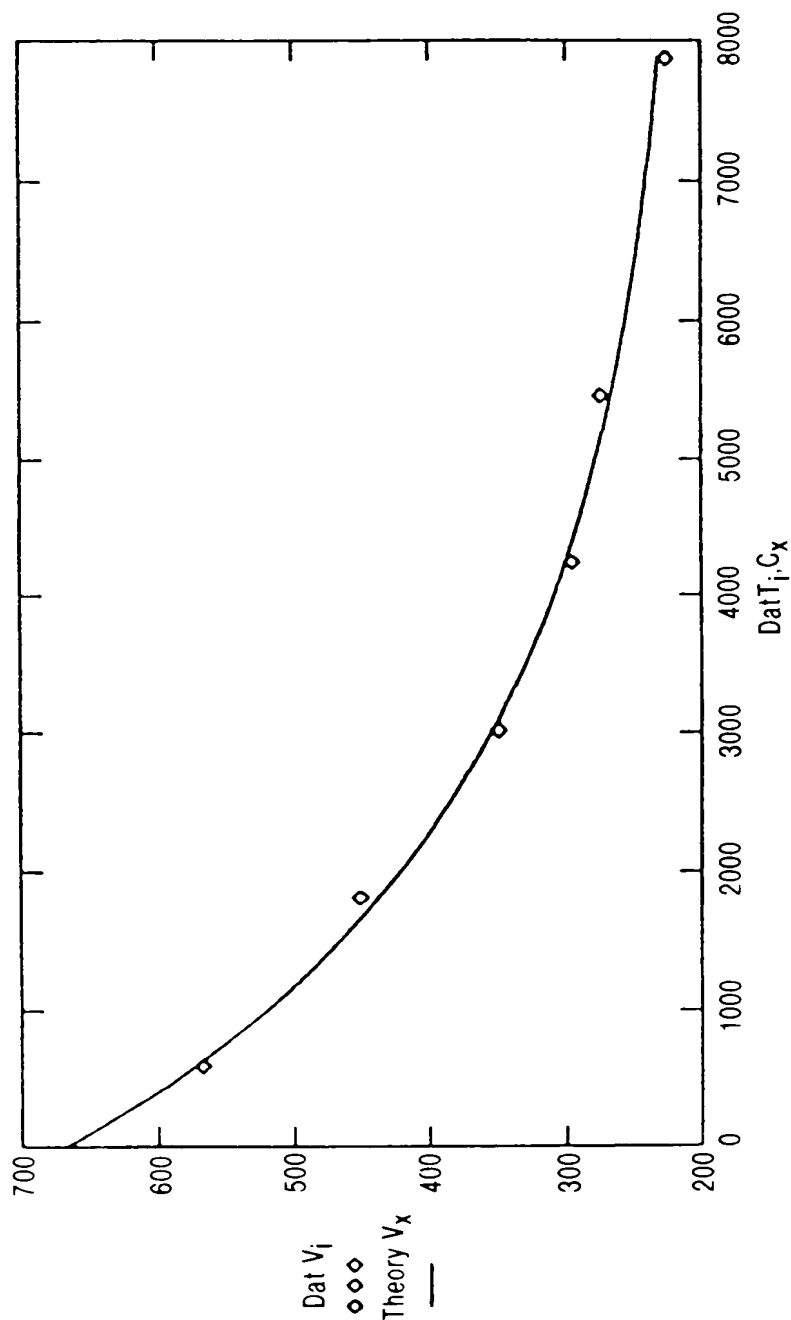


FIG. 50B



Representative Conc versus Time Plot. Shown is the kinetic data for 6G4V11N35A.IgG1

SAMPLE	k_a	k_d	Kd
Murine 6G4.2.5 IgG2a	8.3×10^5	2.9×10^{-4}	350pM
6G4V11N35A-IgG1	8.7×10^5	7.7×10^{-5}	88pM
6G4V11N35E-IgG1	3.0×10^6	1.4×10^{-4}	49pM

FIG. 51

JAN 22 2003

Title: Antibody Fragment-Polymer Conjugates and Humanized Anti-IL-8

Monoclonal Antibodies; Inventor: Vanessa H. et al.

Application No.: 09/234,182 (Attorney Docket No. 09-000000093A)

111/141

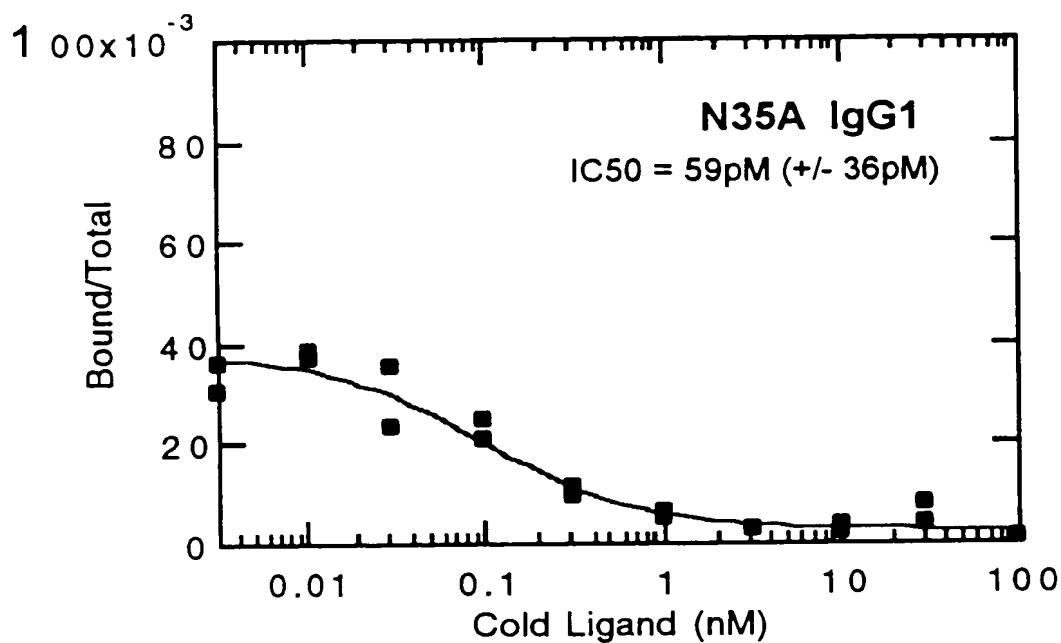


FIG. 52A

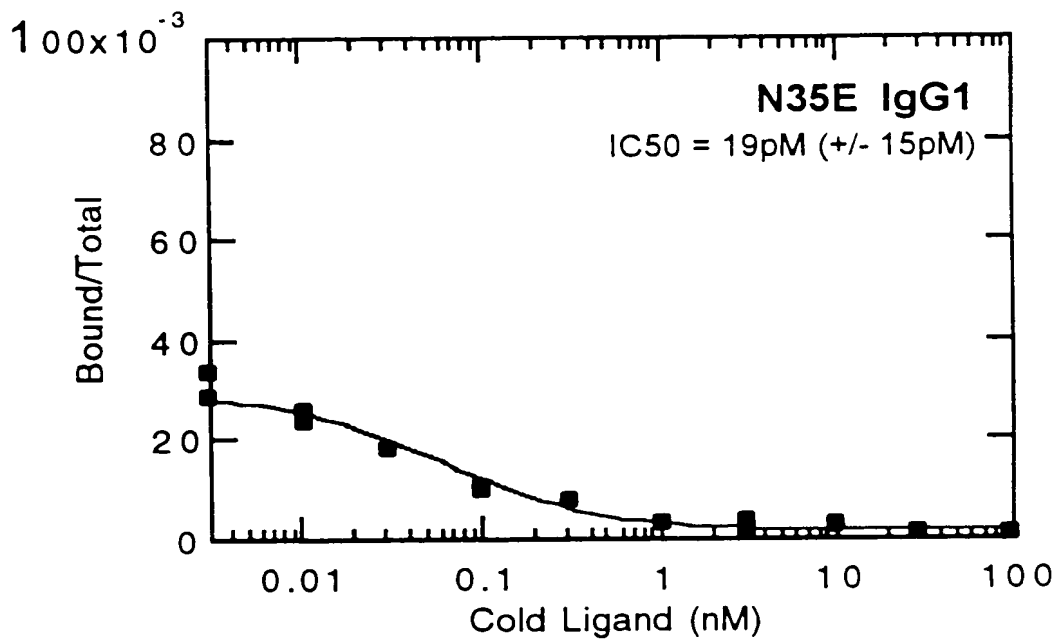


FIG. 52B

781 AAAAGGGTAT CTAGAGGTTG AGGTGATTTT ATGAAAAAGA ATATCGCATT TCTTCTTGCA
 TTTTCCCATA GATCTCCAAC TCCACTAAAA TACTTTTTTCT TATAGCGTAA AGAAGAACGT
 -1 M K K N I A F L L A
 841 TCTATGTTTCG TTTTTTCTAT TGCTACAAAC GCGTACGCTG AGGTTTCAGCT AGTGCAGTCT
 AGATACAAGC AAAAAAGATA ACGATGTTTG CGCATGCGAC TCCAAGTCGA TCACGTCAGA
 -11 S M F V F S I A T N A Y A E V Q L V Q S
 901 GGCGGTGGCC TGGTGCAGCC AGGGGGCTCA CTCCGTTTGT CCTGTGCAGC TTCTGGCTAC
 CCGCCACCGG ACCACGTCGG TCCCCCGAGT GAGGCAAACA GGACACGTCG AAGACCGATG
 8 G G G L V Q P G G S L R L S C A A S G Y
 961 TCCTTCTCGA GTCACTATAT GCACTGGGTC CGTCAGGCCC CGGGTAAGGG CCTGGAATGG
 AGGAAGAGCT CAGTGATATA CGTGACCCAG GCAGTCCGGG GCCCATTCCC GGACCTTACC
 28 S F S S H Y M H W V R Q A P G K G L E W
 1021 GTTGGATATA TTGATCCTTC CAATGGTGAA ACTACGTATA ATCAAAAGTT CAAGGGCCGT
 CAACCTATAT AACTAGGAAG GTTACCACCT TGATGCATAT TAGTTTTCAA GTTCCCGGCA
 48 V G Y I D P S N G E T T Y N Q K F K G R
 1081 TTCACCTTTAT CTCGCGACAA CTCCAAAAAC ACAGCATACC TGCAGATGAA CAGCCTGCGT
 AAGTGAAATA GAGCGCTGTT GAGGTTTTTG TGTCGTATGG ACGTCTACTT GTCGGACGCA
 68 F T L S R D N S K N T A Y L Q M N S L R
 1141 GCTGAGGACA CTGCCGTCTA TTA CTGTGCA AGAGGGGATT ATCGCTACAA TGGTGA CTGG
 CGACTCCTGT GACGGCAGAT AATGACACGT TCTCCCCCTAA TAGCGATGTT ACCACTGACC
 88 A E D T A V Y Y C A R G D Y R Y N G D W
 1201 TTCTTCGACG TCTGGGGTCA AGGAACCCTG GTCACCGTCT CCTCGGCCTC CACCAAGGGC
 AAGAAGCTGC AGACCCCACT TCCTTGGGAC CAGTGCCAGA GGAGCCGGAG GTGGTTCCCG
 108 F F D V W G Q G T L V T V S S A S T K G
 1261 CCATCGGTCT TCCCCCTGGC ACCCTCCTCC AAGAGCACCT CTGGGGGCAC AGCGGCCCTG
 GG TAGCCAGA AGGGGGACCG TGGGAGGAGG TTCTCGTGGA GACCCCGTG TCGCCGGGAC
 128 P S V F P L A P S S K S T S G G T A A L
 1321 GGCTGCCTGG TCAAGGACTA CTTCCCCGAA CCGGTGACGG TGTCGTGGAA CTCAGGCGCC
 CCGACGGACC AGTTCTTGAT GAAGGGGCTT GGCCACTGCC ACAGCACCTT GAGTCCGCGG
 148 G C L V K D Y F P E P V T V S W N S G A
 1381 CTGACCAGCG GCGTGACAC CTTCCCGGCT GTCCTACAGT CCTCAGGACT CTACTCCCTC
 GACTGGTTCG CGCACGTGTG GAAGGGCCGA CAGGATGTCA GGAGTCCTGA GATGAGGGAG
 168 L T S G V H T F P A V L Q S S G L Y S L
 1441 AGCAGCGTGG TGACCGTGCC CTCCAGCAGC TTGGGCACCC AGACCTACAT CTGCAACGTG
 TCGTCGCACC ACTGGCACGG GAGGTCGTG AACCCGTGGG TCTGGATGTA GACGTTGCAC
 188 S S V V T V P S S S L G T Q T Y I C N V
 1501 AATCACAAGC CCAGCAACAC CAAGGTCGAC AAGAAAGTTG AGCCCAAATC TTGTGACAAA
 TTAGTGTTTCG GGTGTTGTG GTTCCAGCTG TTCTTTCAAC TCGGGTTTAG AACACTGTTT
 208 N H K P S N T K V D K K V E P K S C D K
 1561 ACTCACACAT GCGCCGCGTGA (SEQ ID NO.69)
 TGAGTGTGTA CGGGCGGCACT
 228 T H T C P P O (SEQ ID NO.70)

FIG. 53

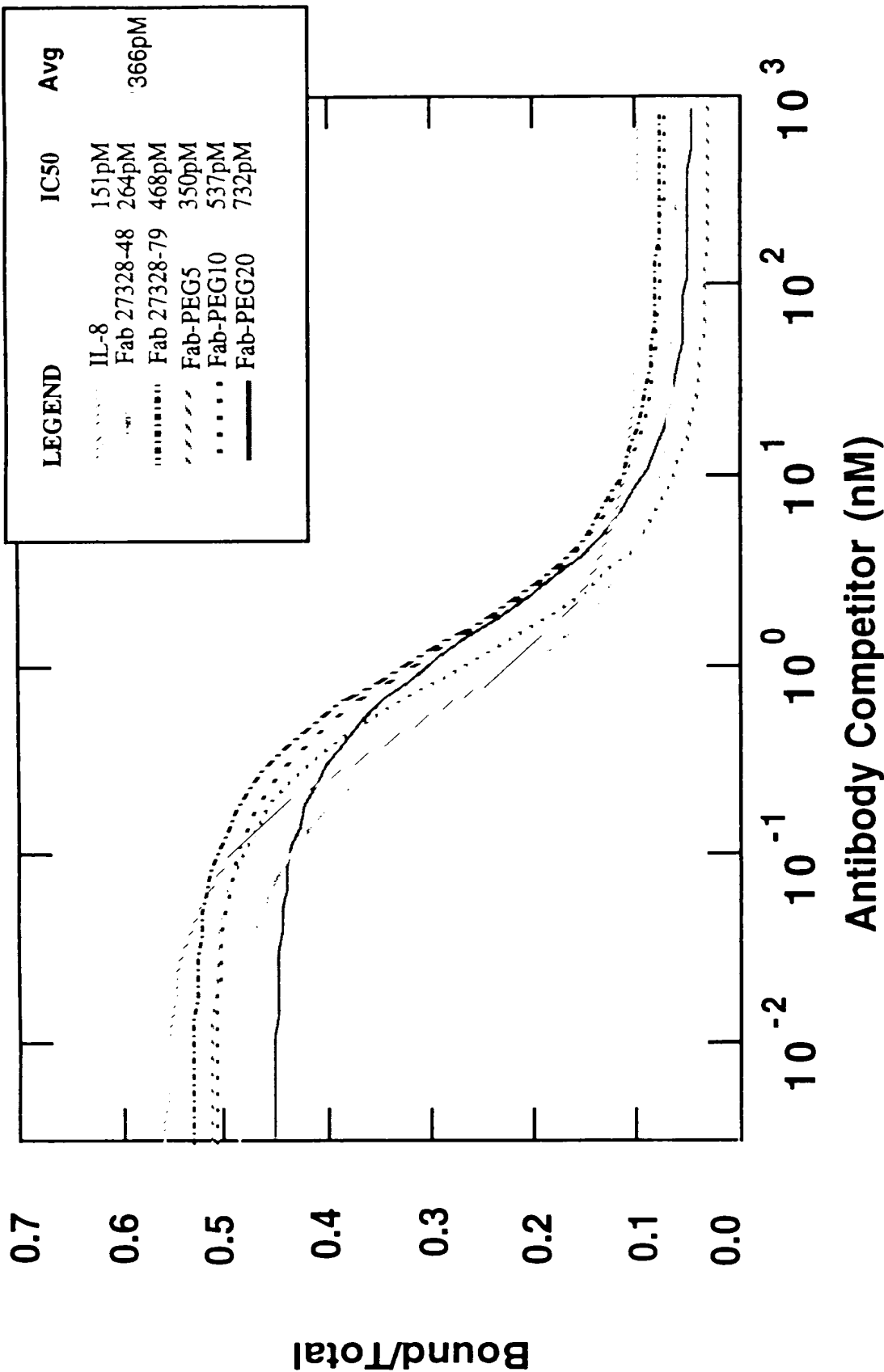
JAN 7 2003

Title: Antibody Fragment-Polymer Conjugates and Humanized Anti-IL-8

Monoclonal Antibodies; Inventor: Vanessa H. et al.

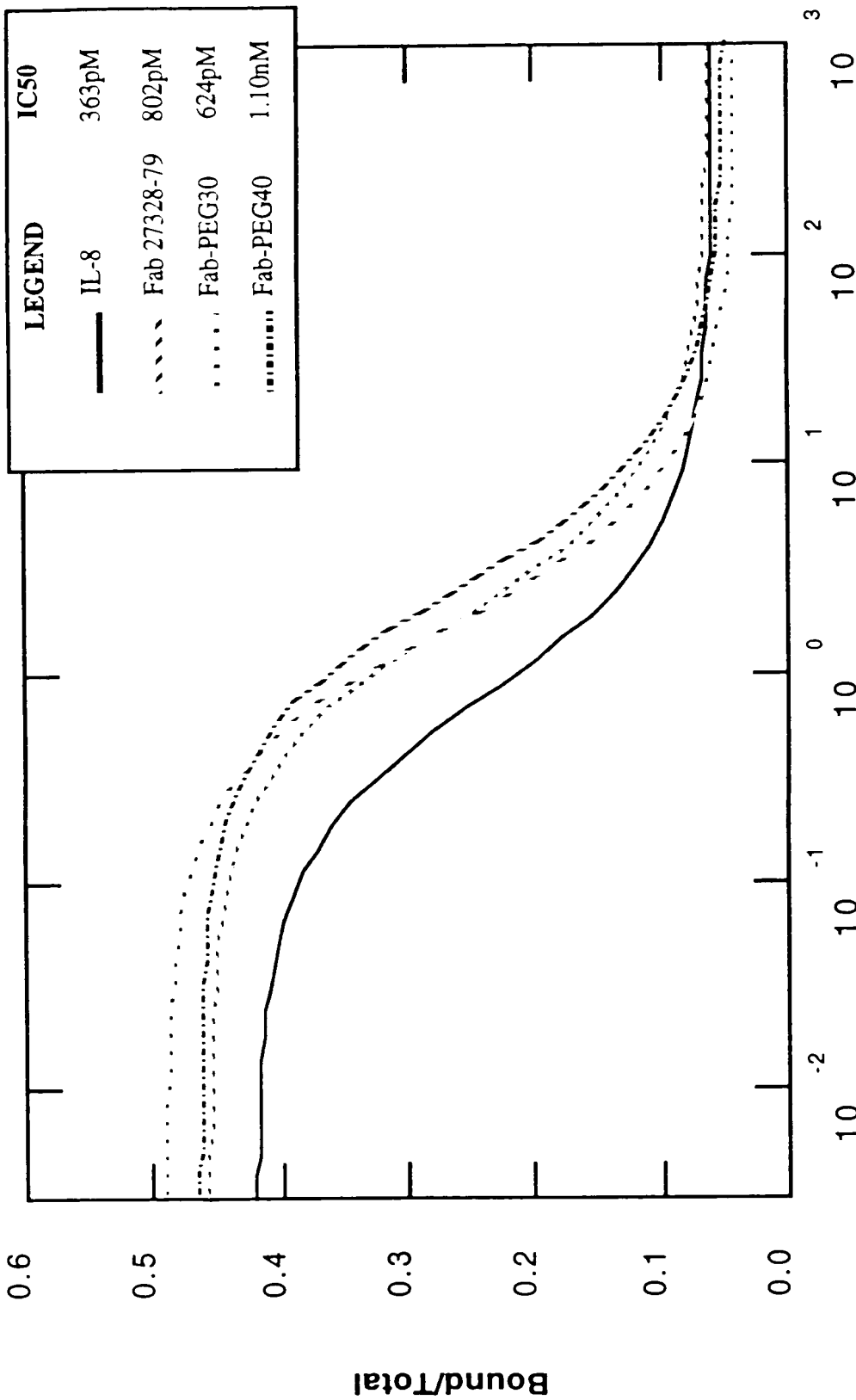
Application No.: 09 234,182 (Attorney Docket No. G000000093A)

113 141



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 JAN 12 2003

BY: [Signature]
 [Signature]
 [Signature]



Antibody Competitor (nM)

FIG. 54B

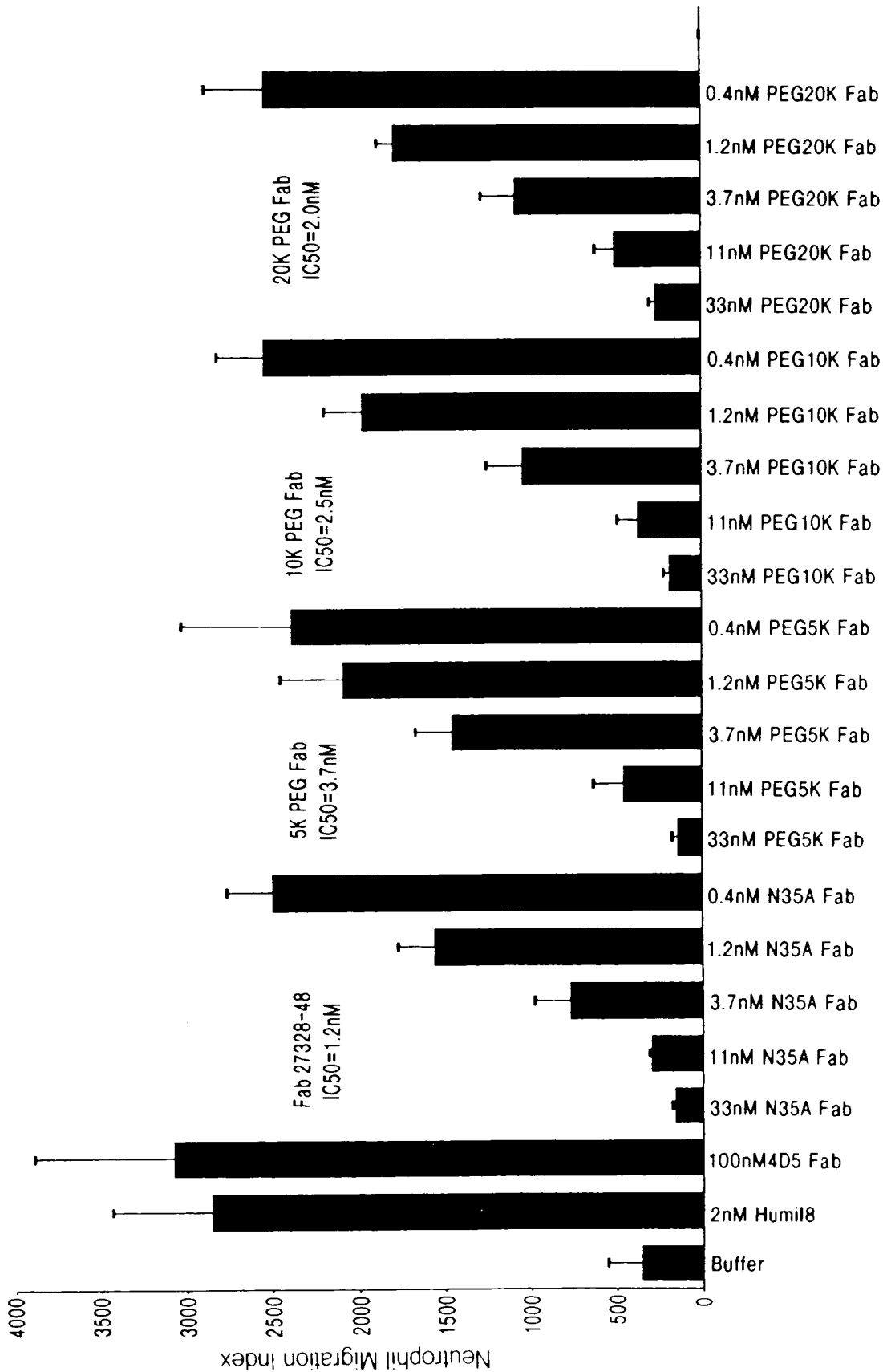
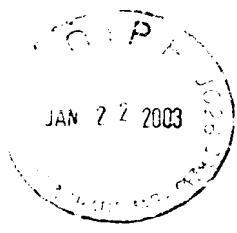
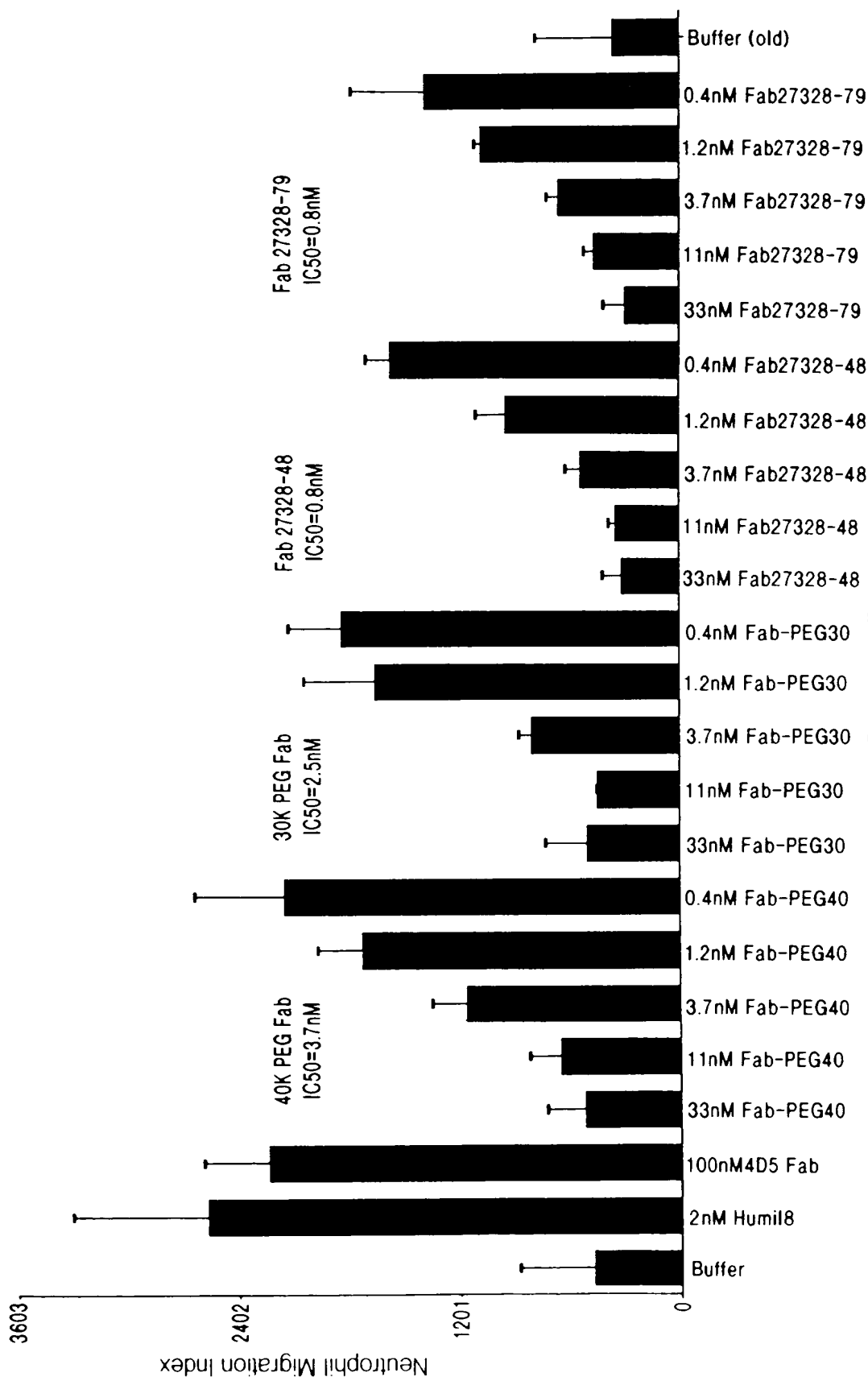
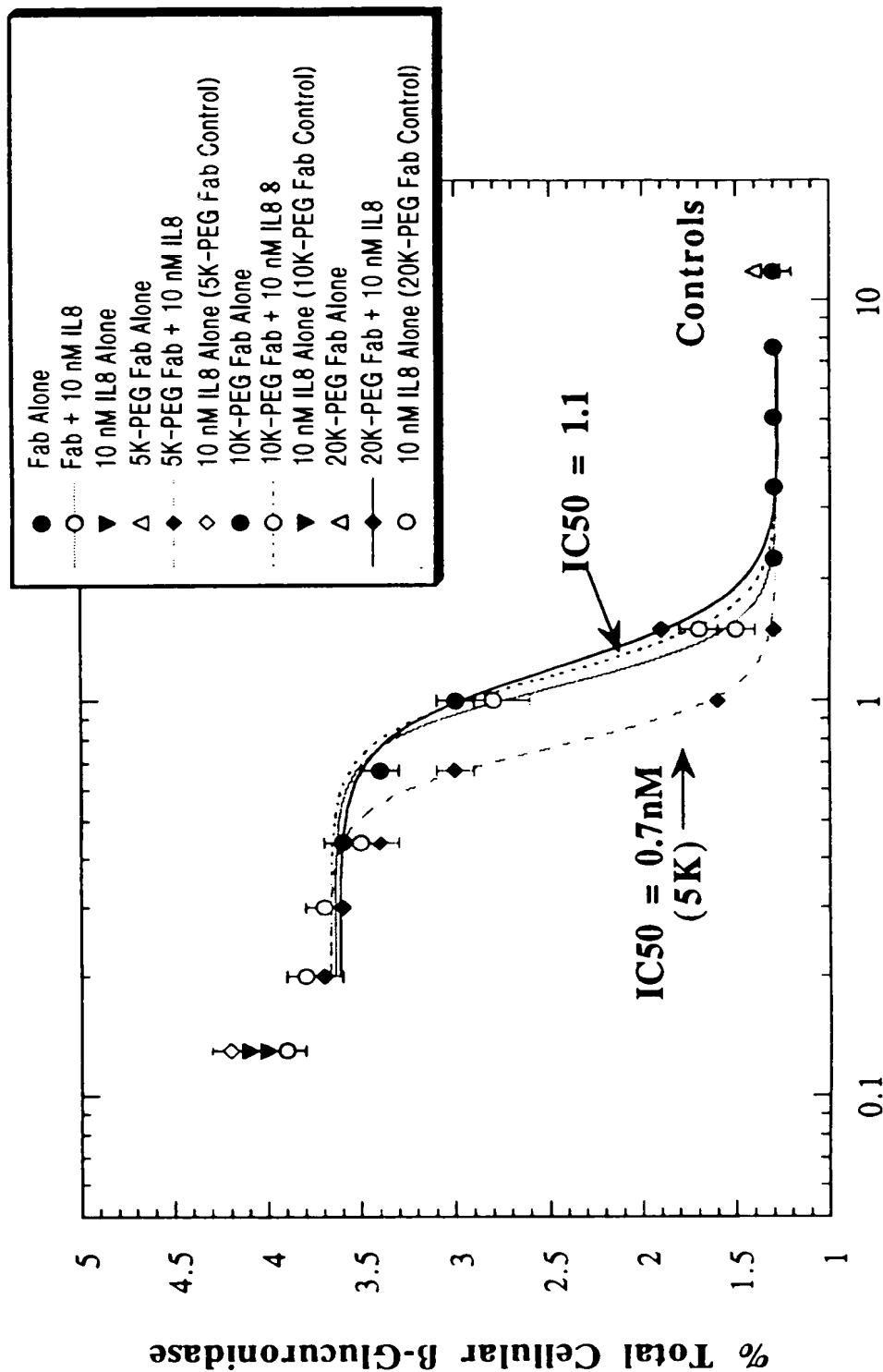


FIG. 55A





Molar Ratio Antibody:IL-8

FIG. 56A



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BY	CLASS	DATE
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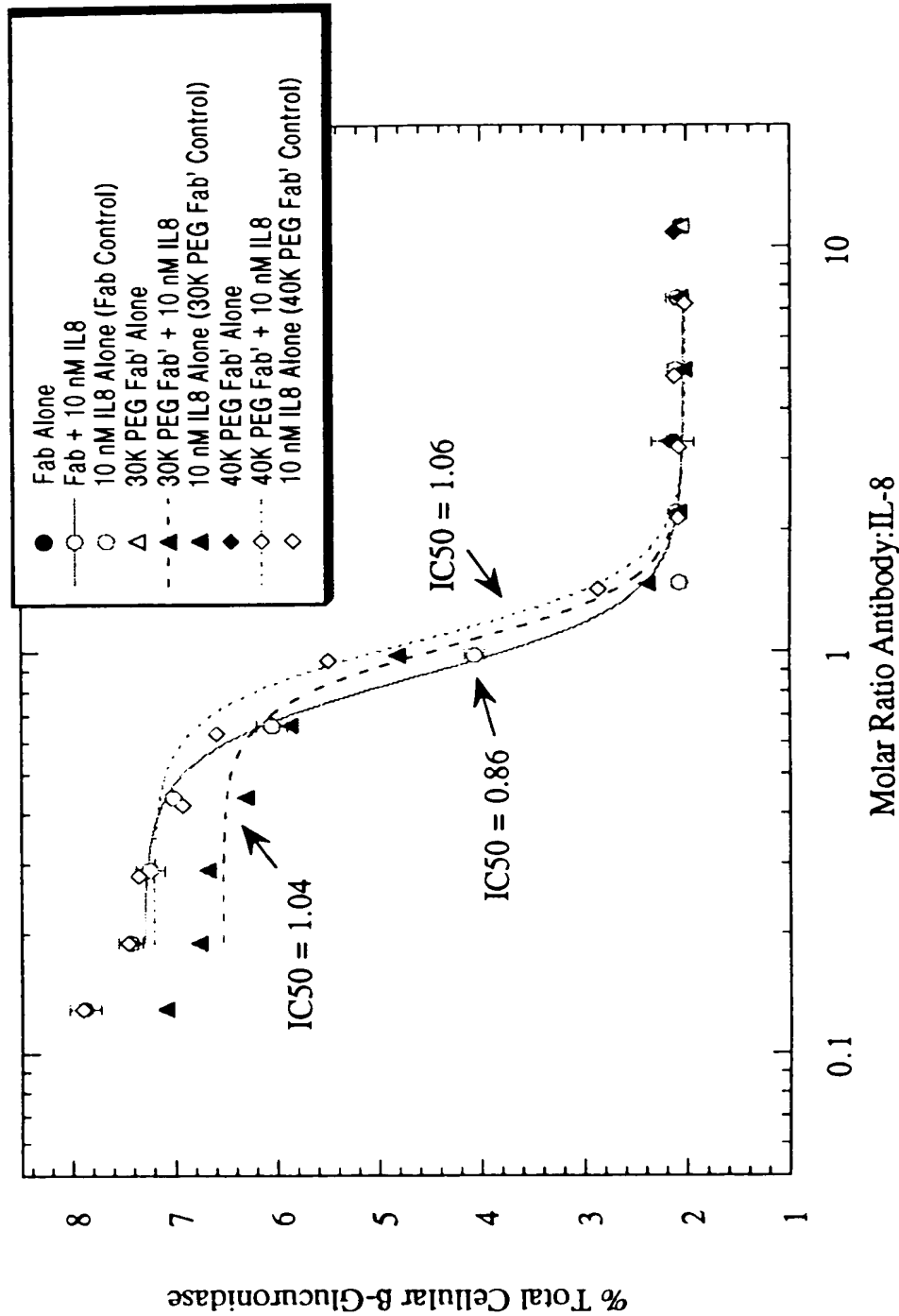
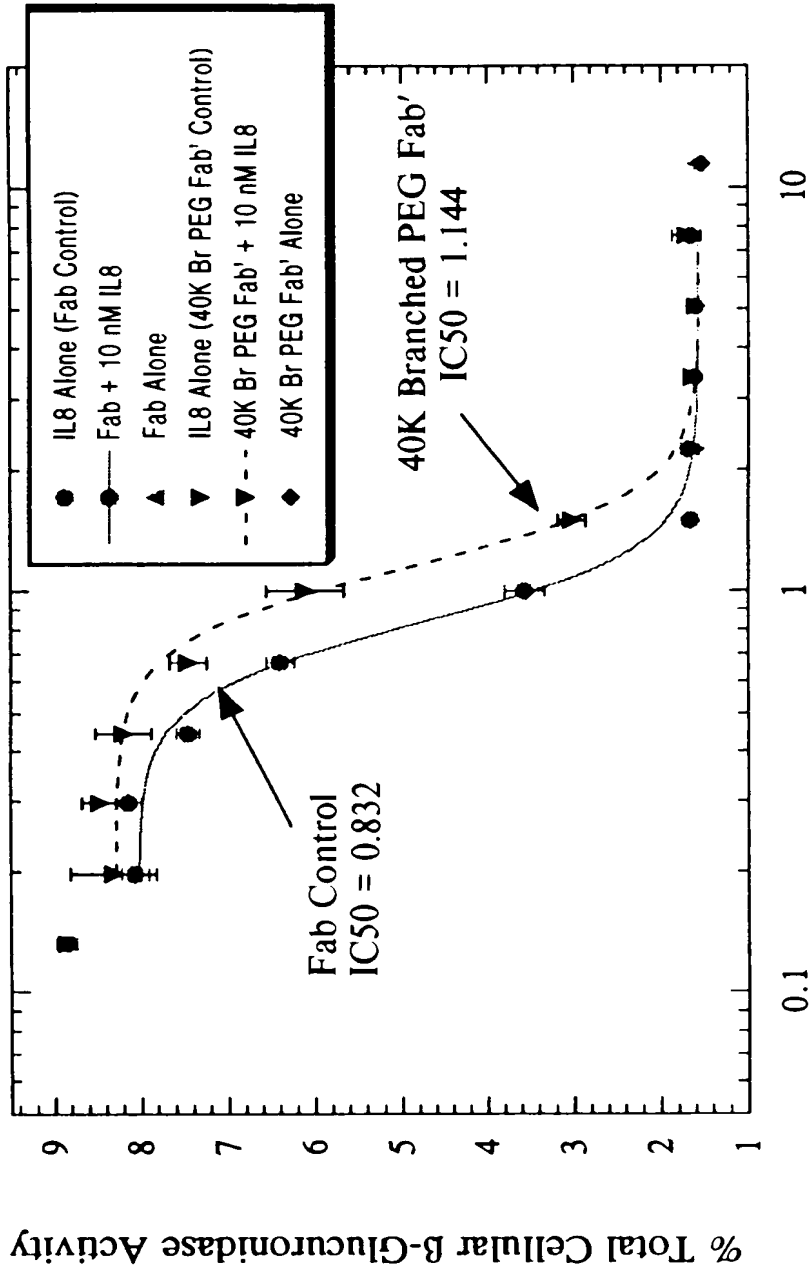


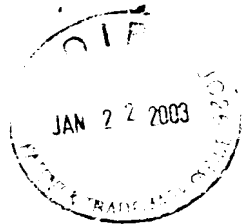
FIG. 56B

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 JAN 27 2003
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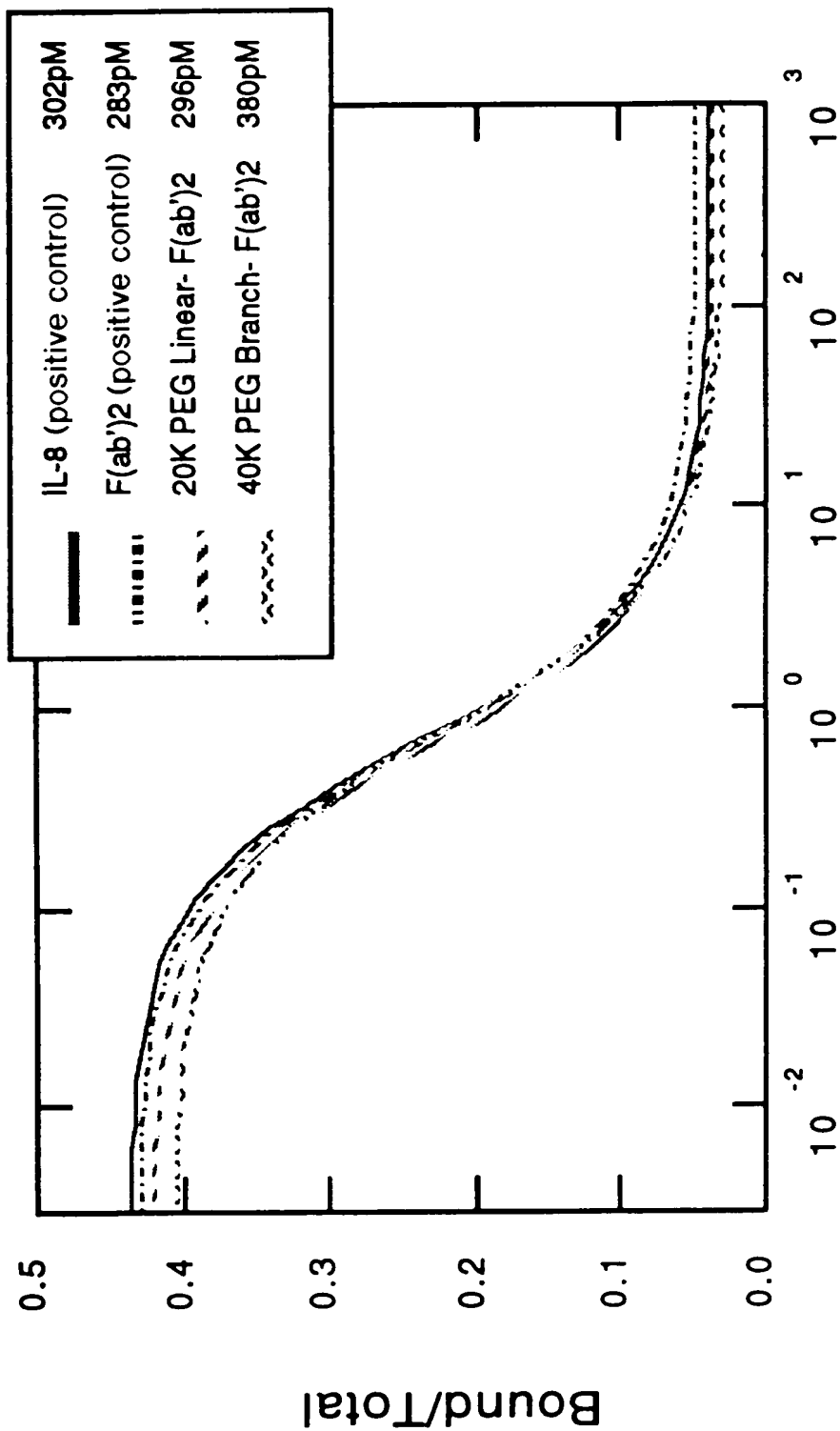


Molar Ratio Antibody:IL8

FIG. 56C

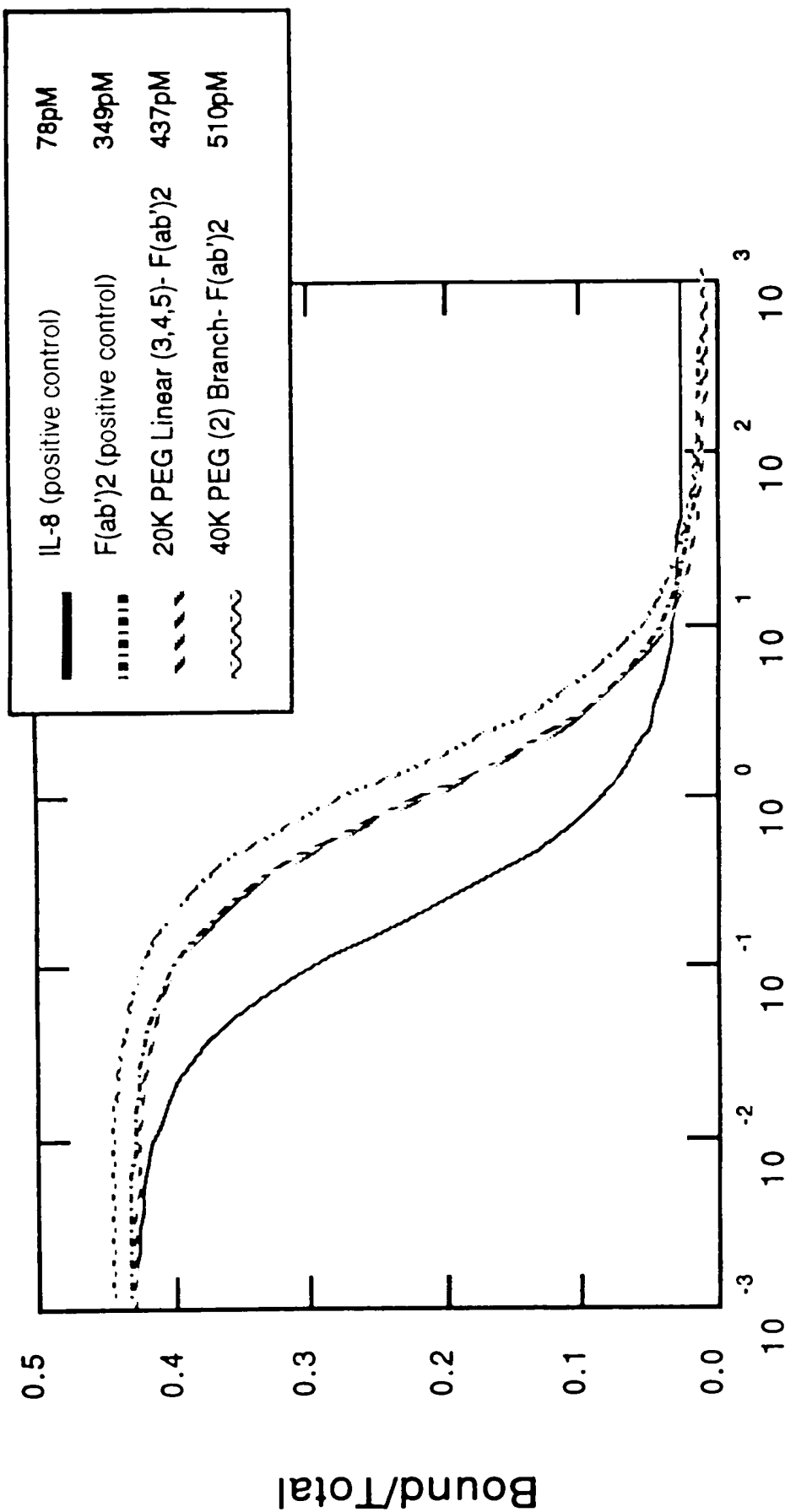
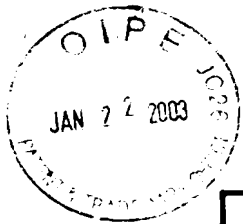


APPROVED	DATE
BY	



Pegylated F(ab')₂ (nM)

FIG. 57A



Pegylated F(ab')₂ (nM)

FIG. 57B

OIPF
 JAN 27 2003
 123 141

Title: Antibody Fragment-Polymer Conjugates and Humanized Anti-IL-8

Monoclonal Antibodies; Inventor: Vanessa H. et al.

Application No.: 09 234,182 (Attorney Docket No. 09234.093A)

123 141

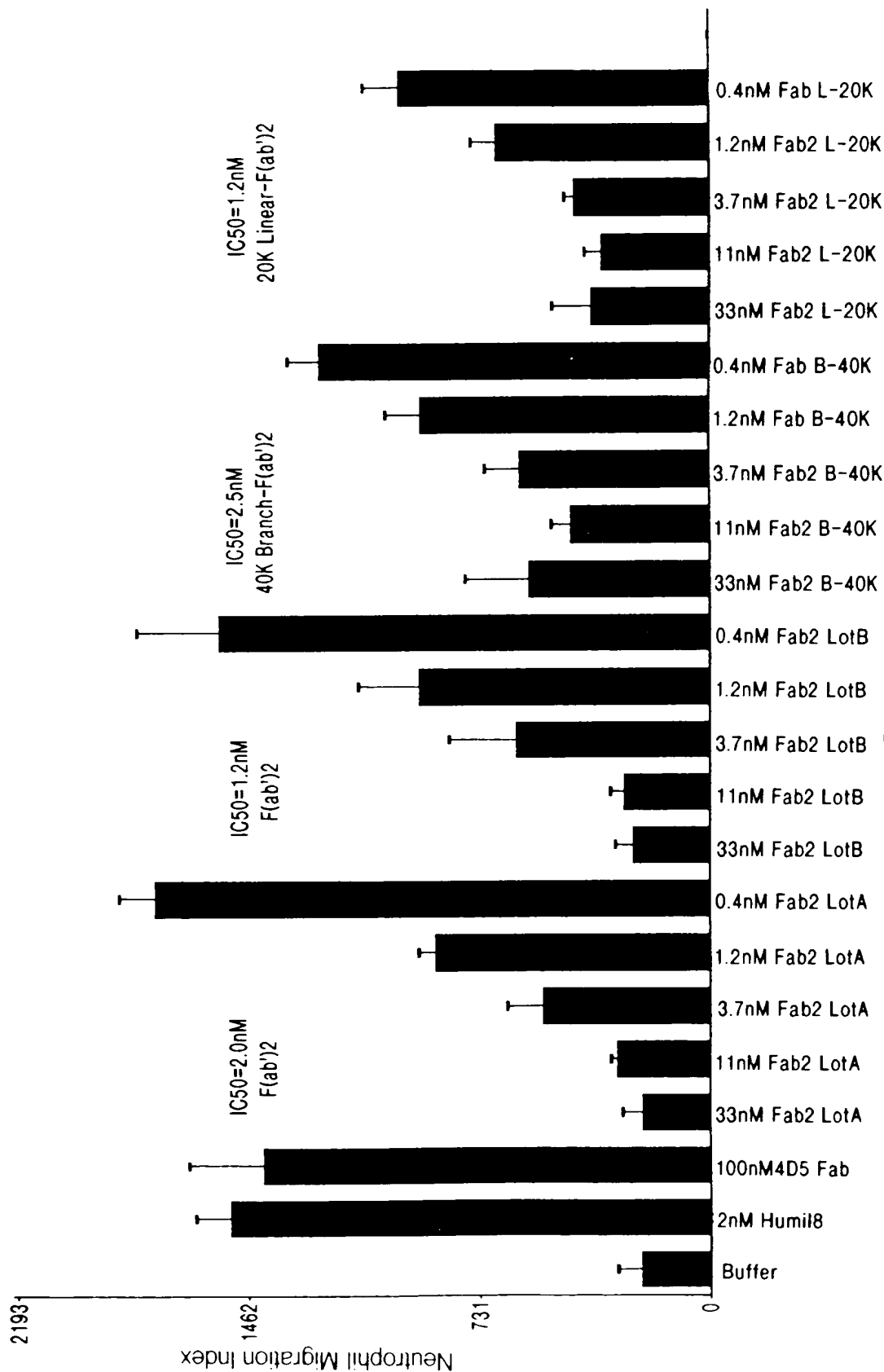


FIG. 58A

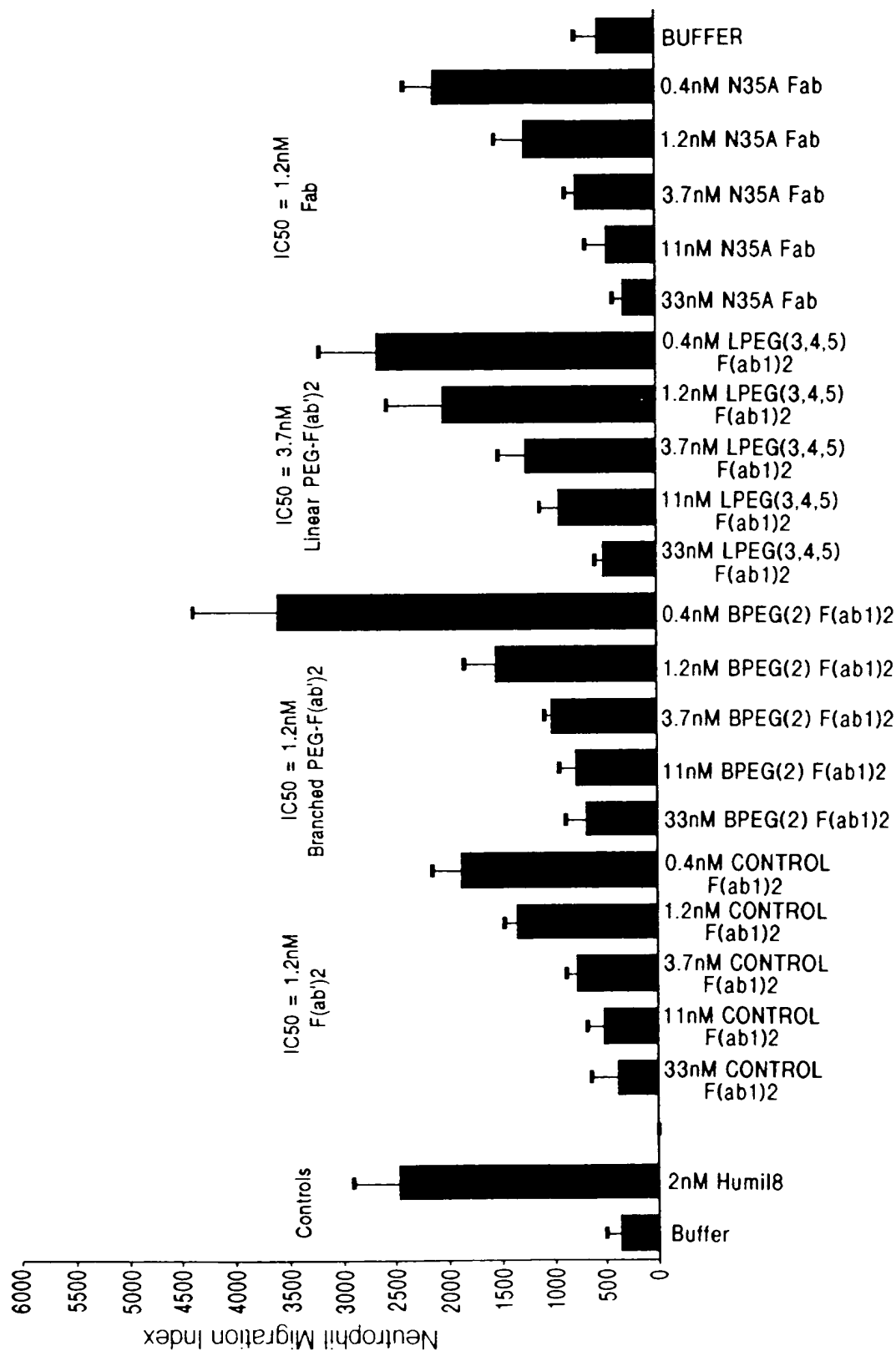


FIG. 58B

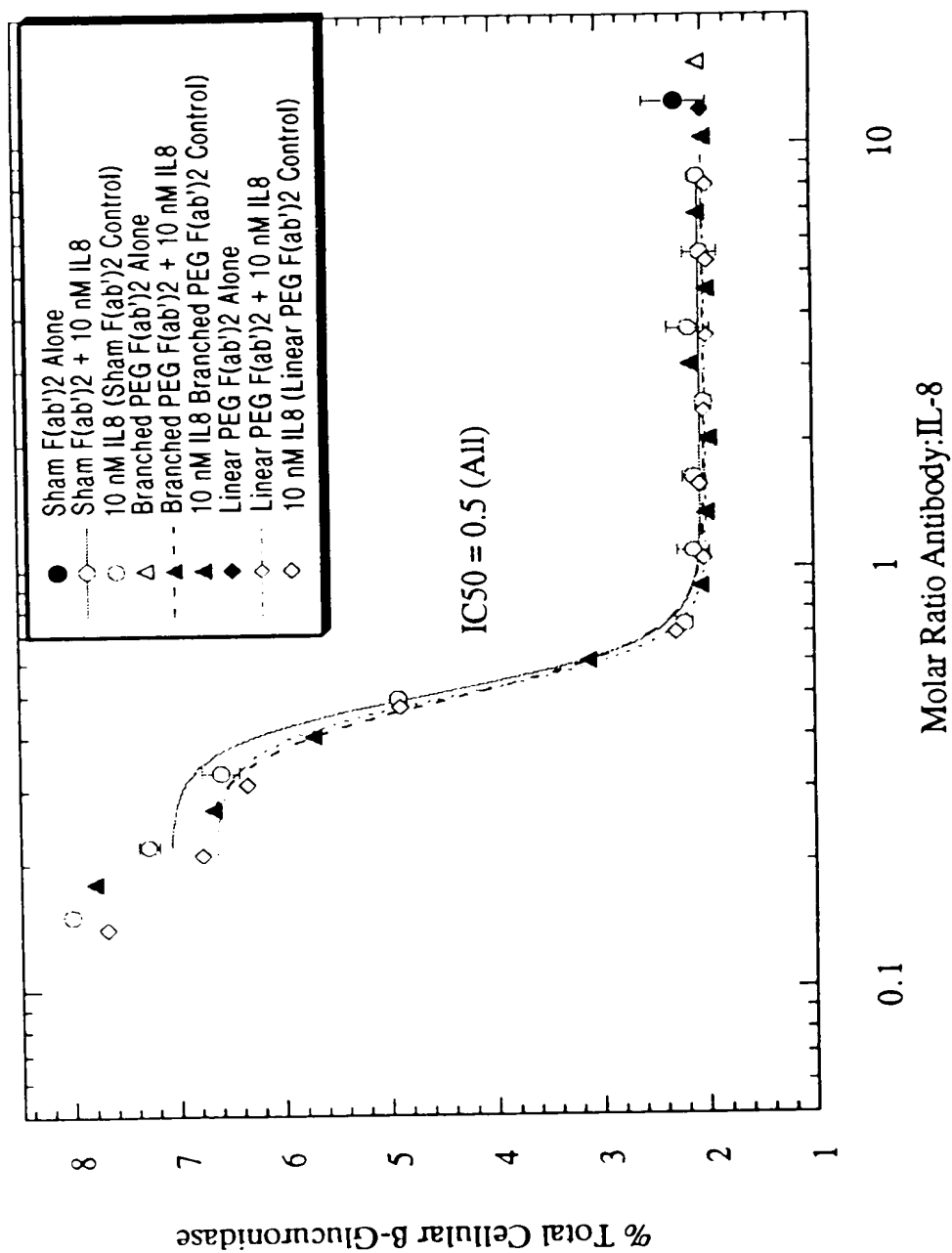


FIG. 59A

0175
JAN 27 2003

Title: Antibody Fragment-Polymer Conjugates and Humanized Anti-IL-8

Monoclonal Antibodies; Inventor: Vanessa F. et al.

Application No.: 09 234,182 (Attorney Docket No. GENENT.093A)

126 141

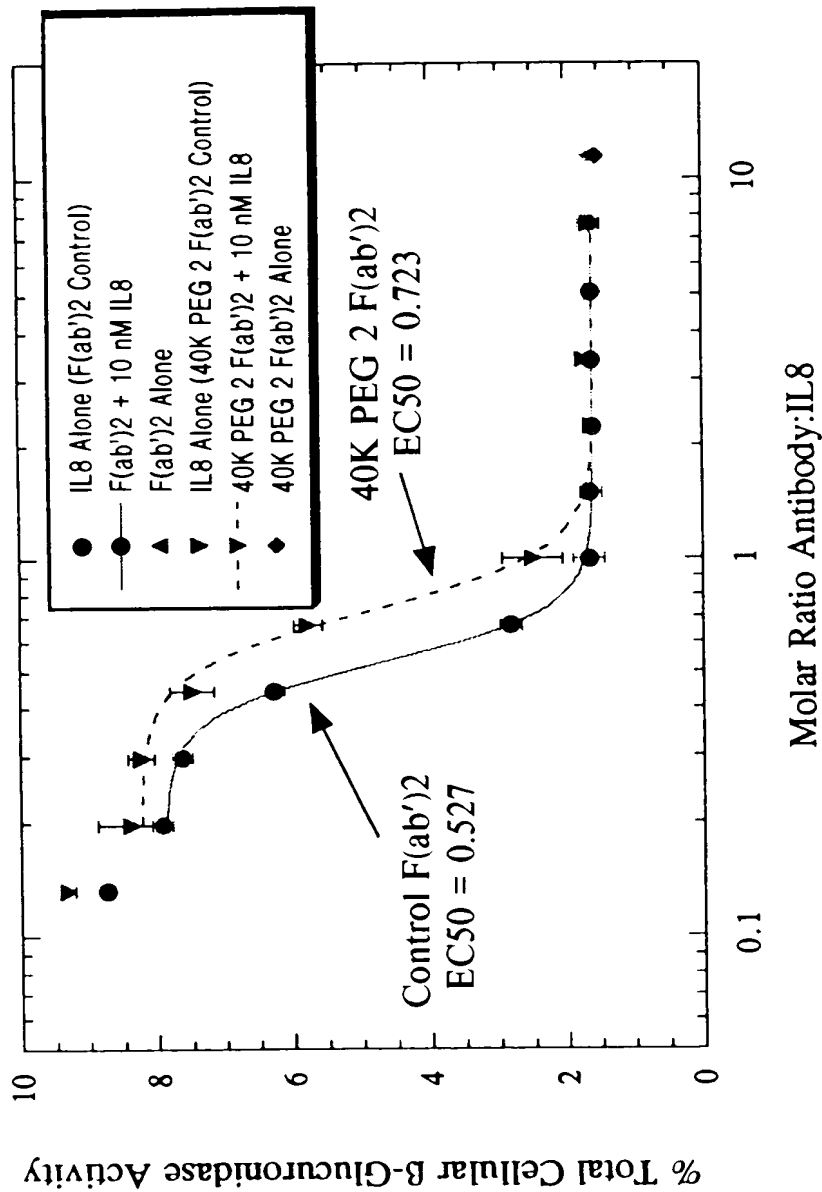
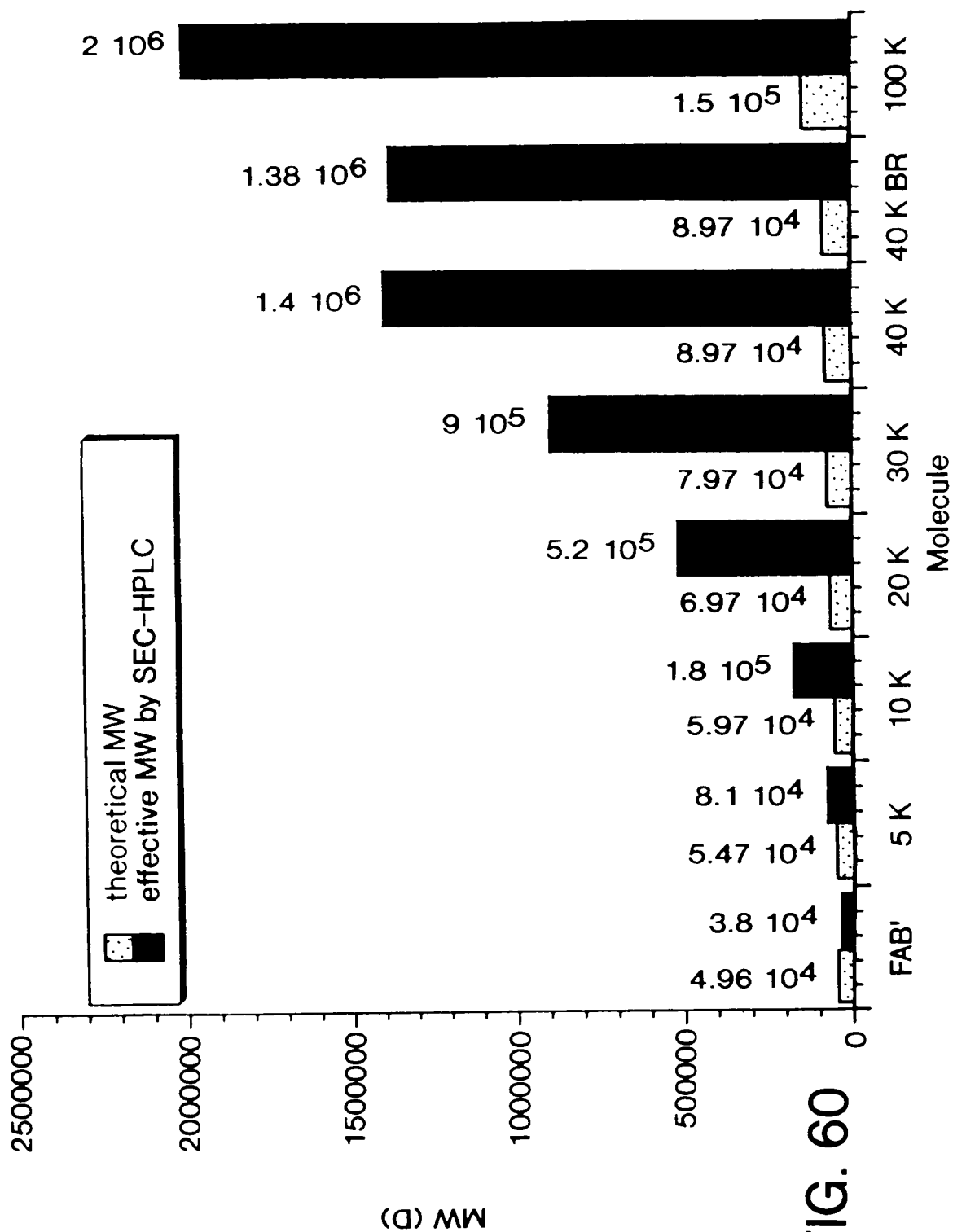
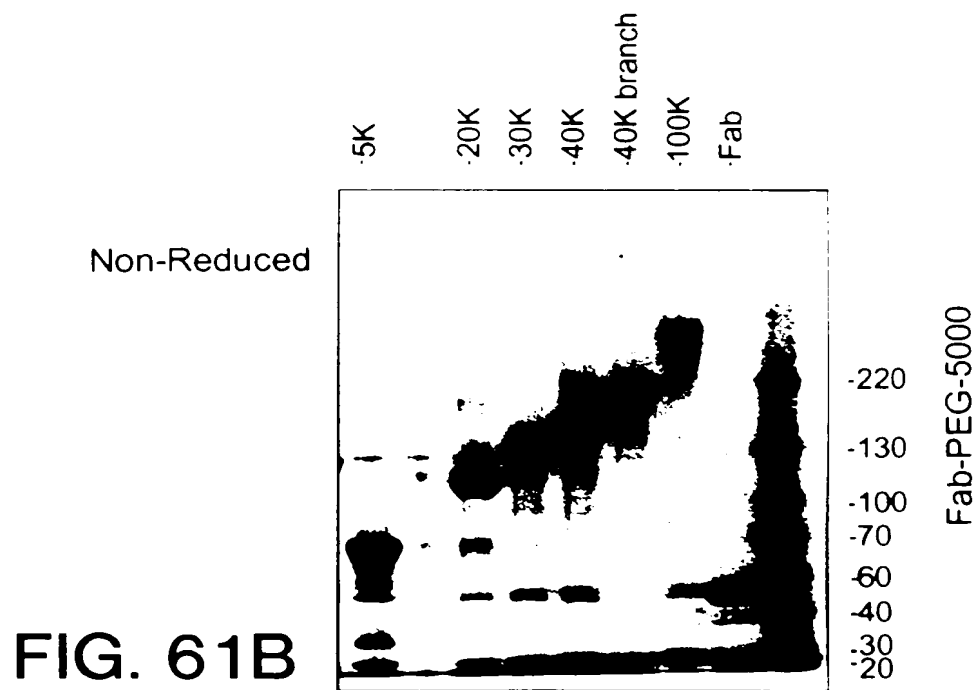
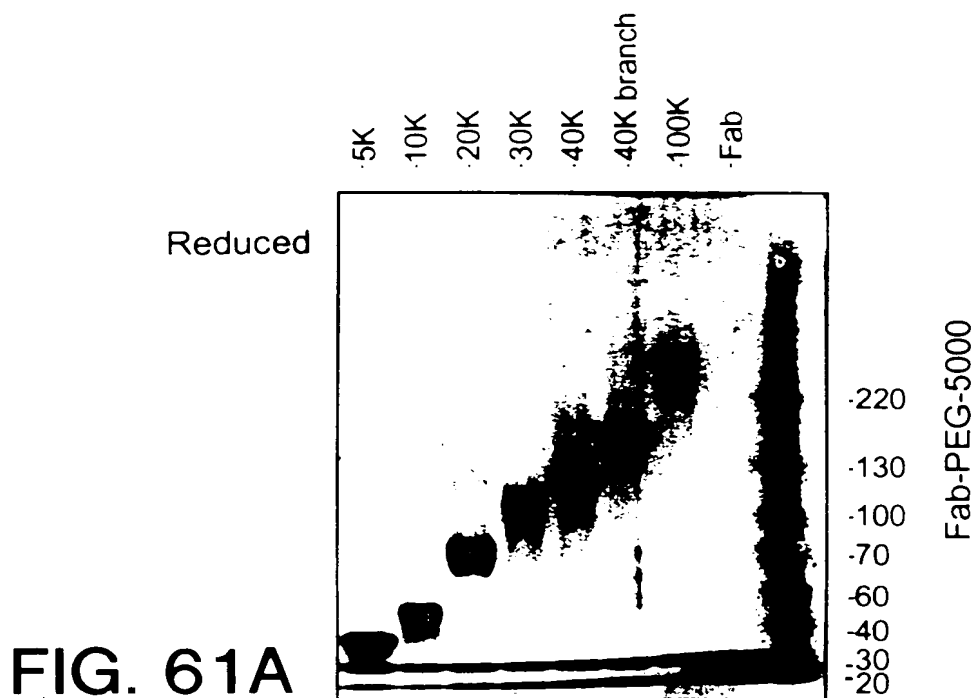


FIG. 59B

JAN 7 2 2003





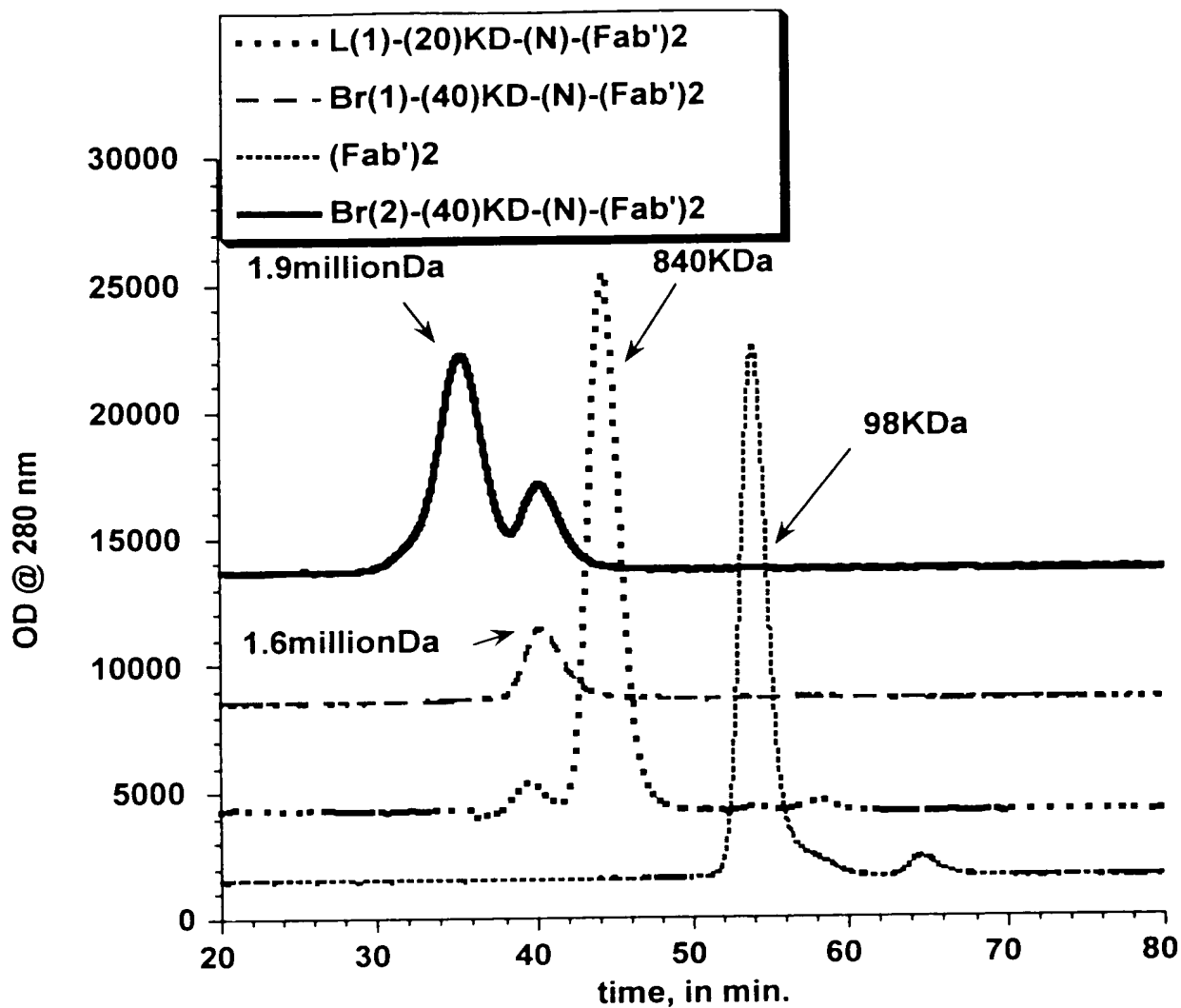


FIG. 62

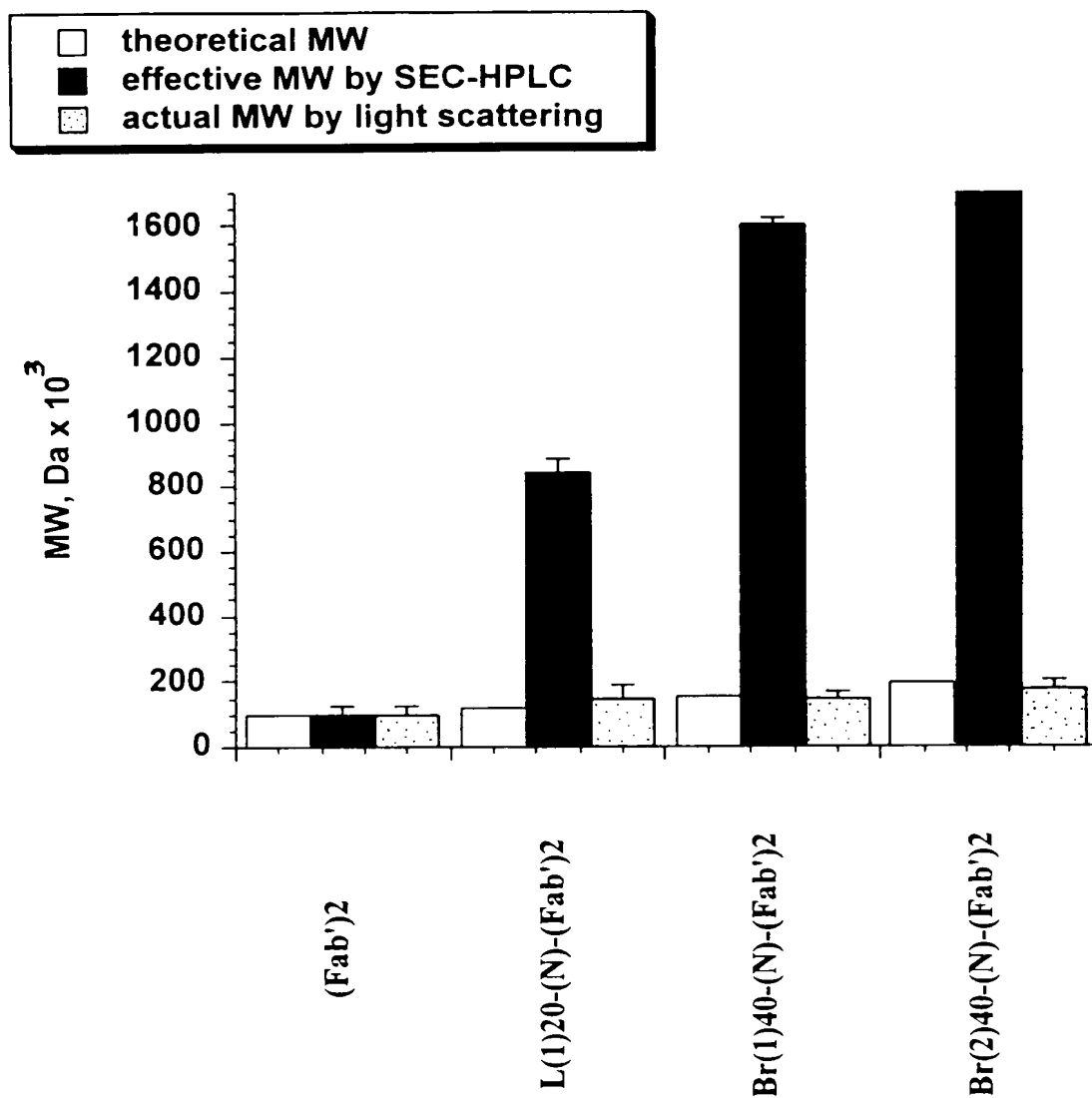


FIG. 63

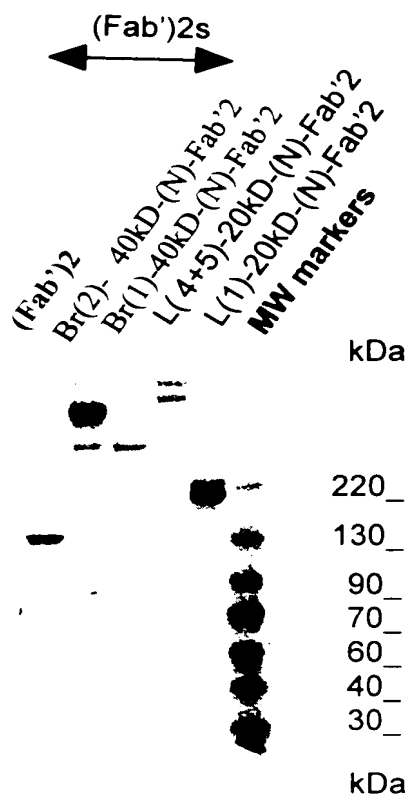
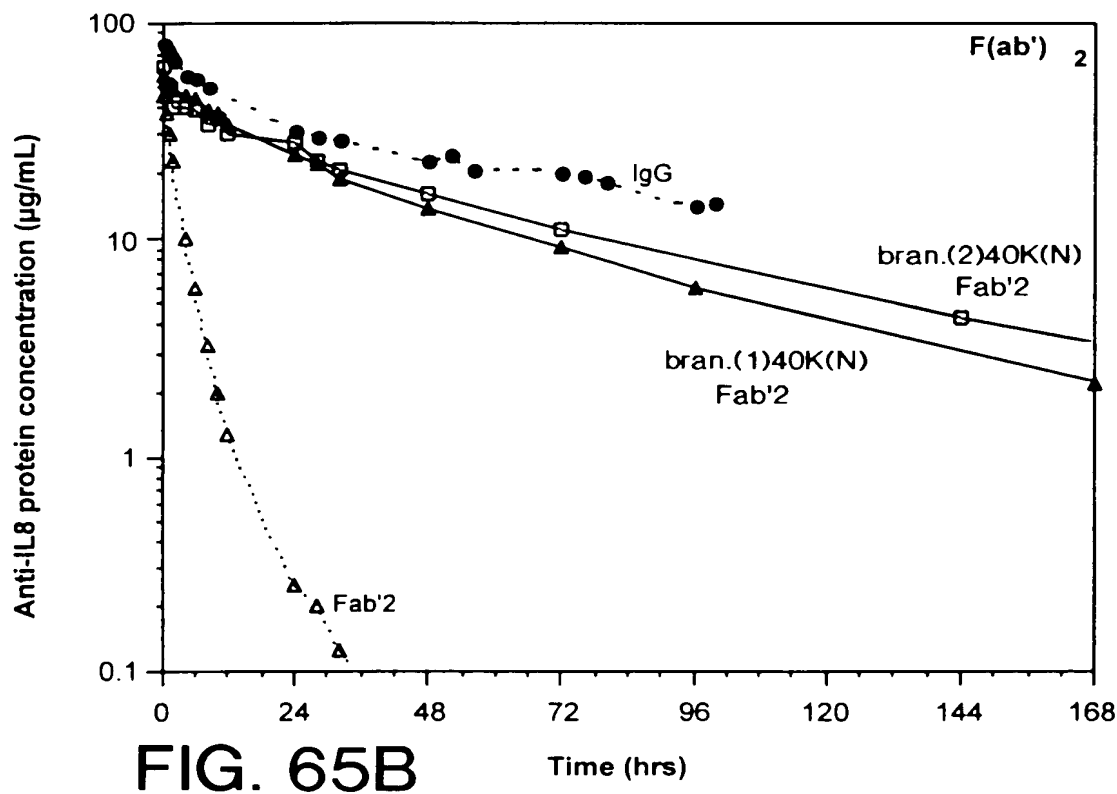
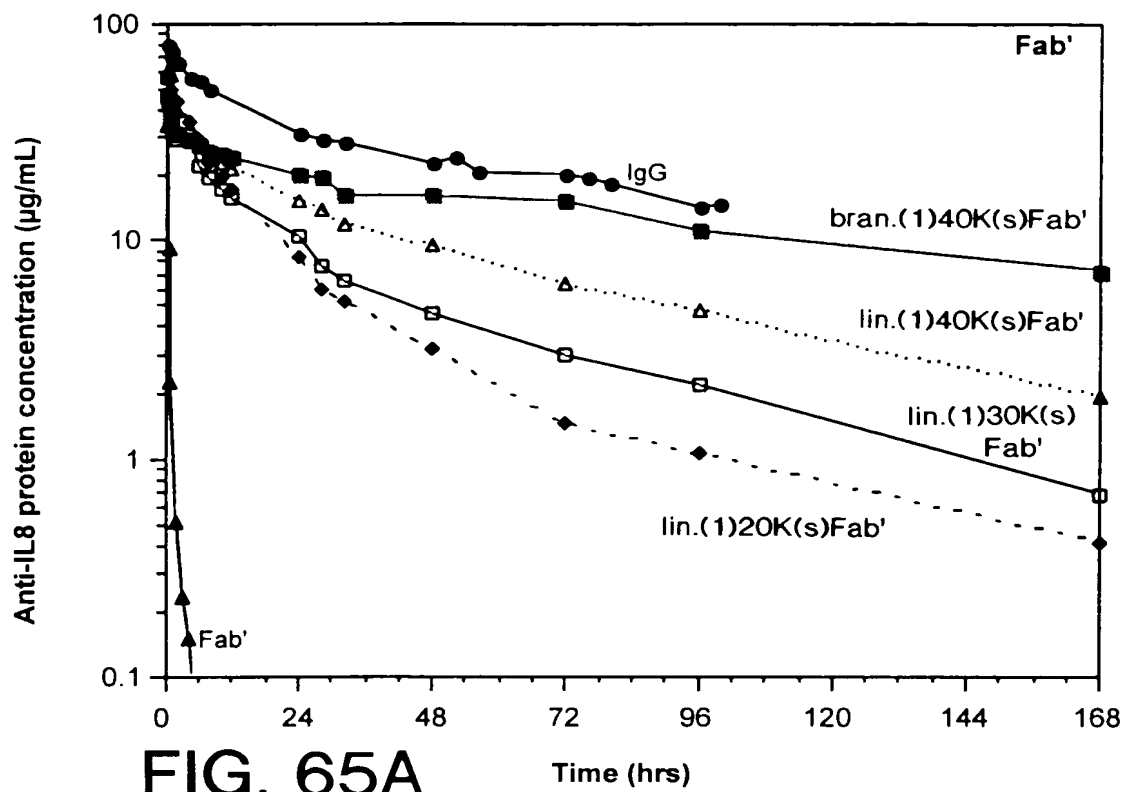


FIG. 64

JAN 7 2003



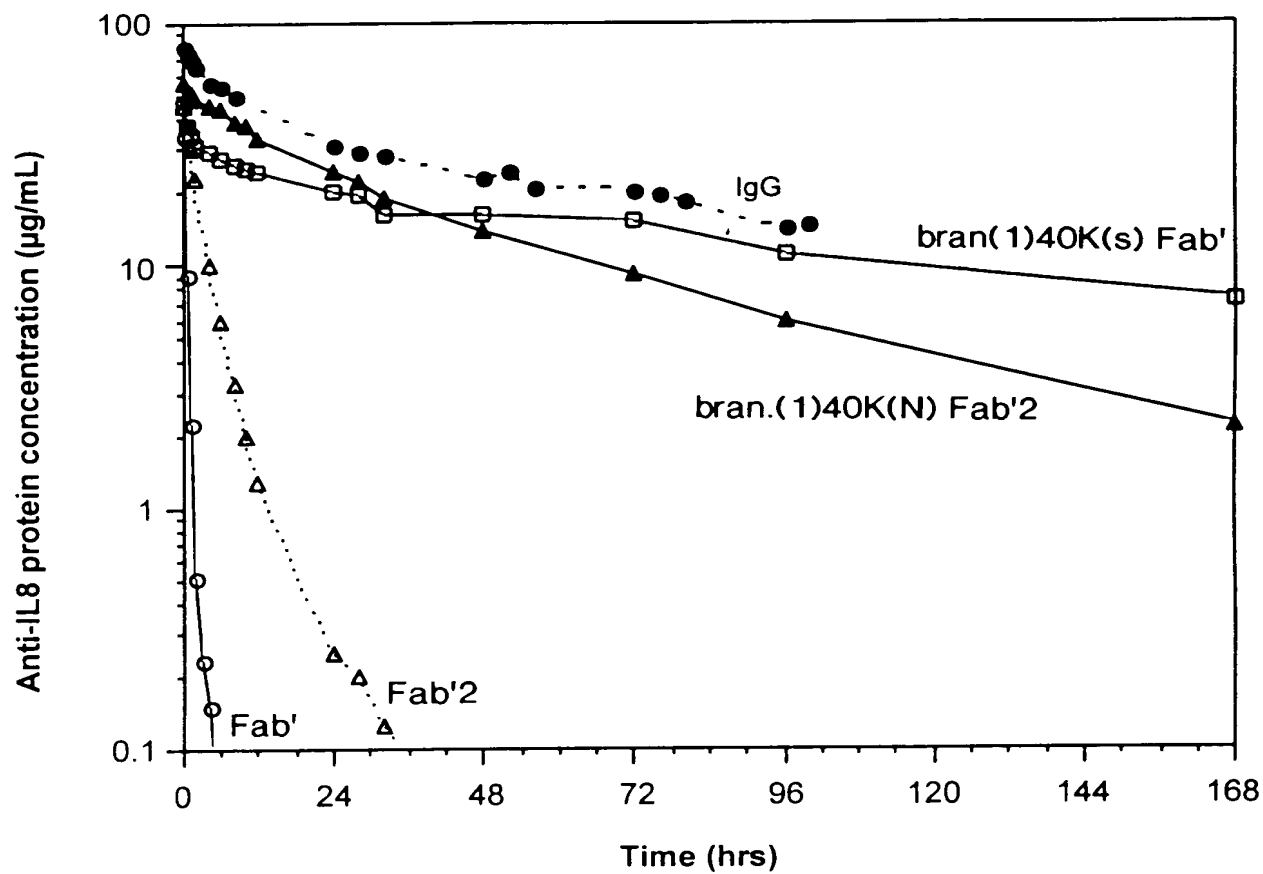


FIG. 66

JAN 27 2003

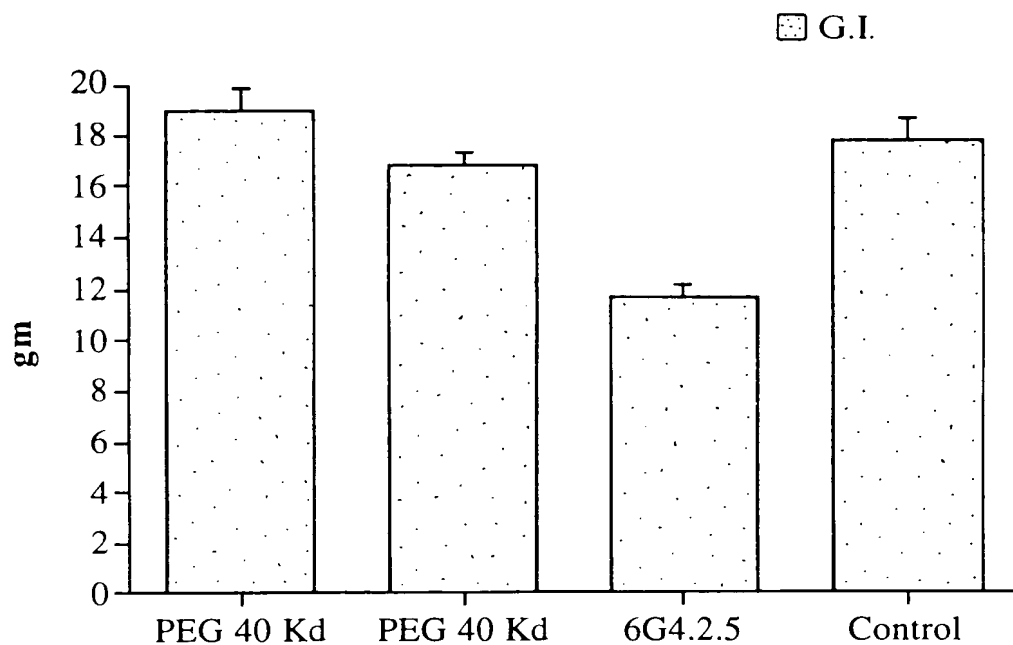


FIG. 67

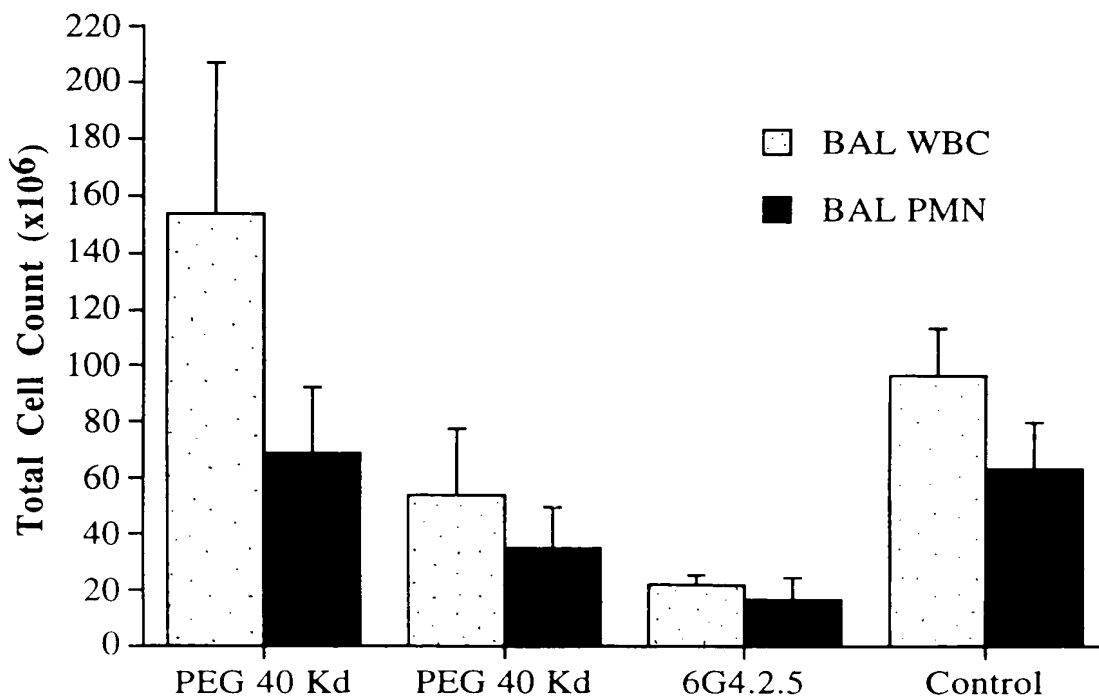


FIG. 68

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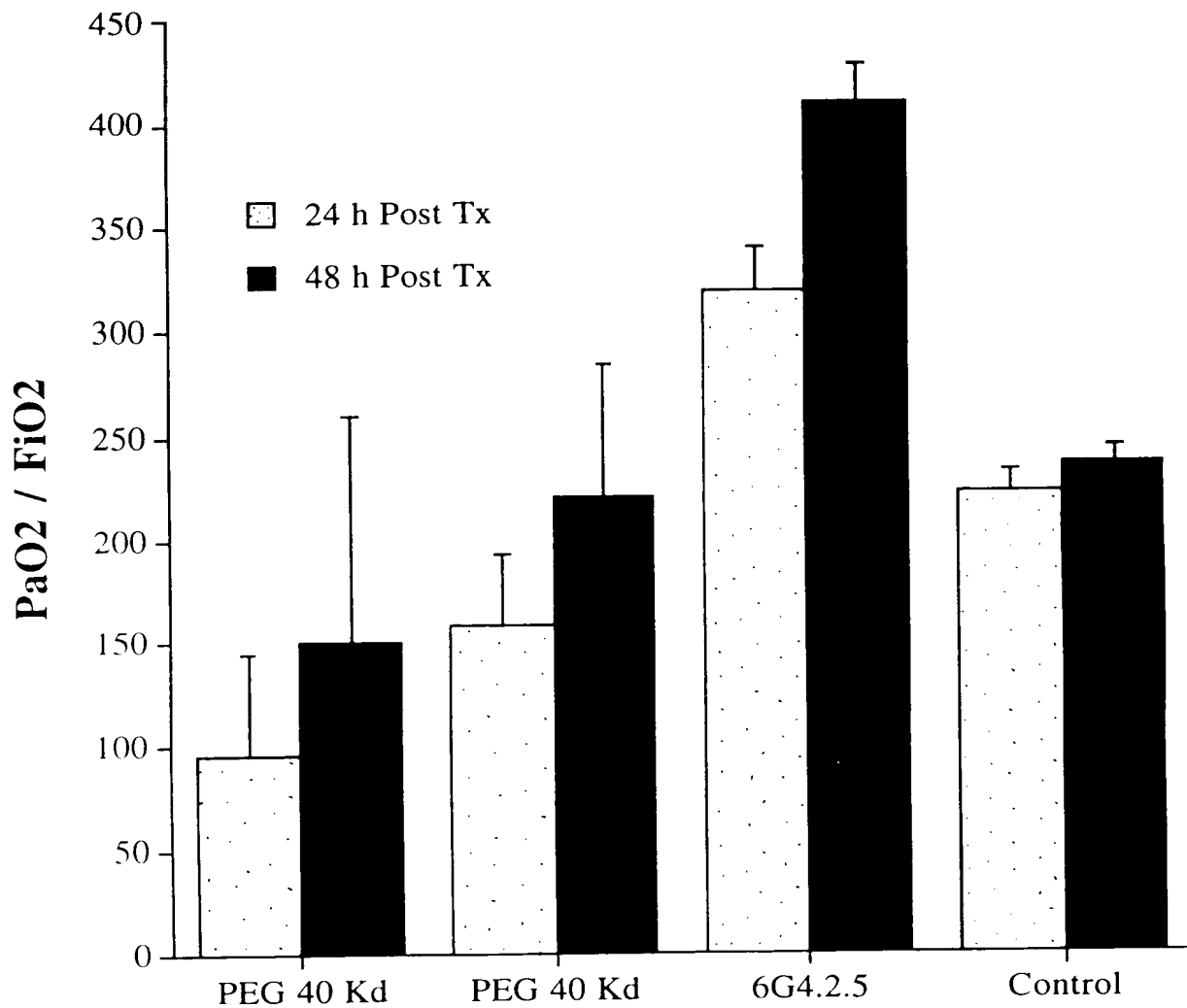


FIG. 69

Oxygenation in 100% O₂ @24 h Post Anti-IL8 Tx

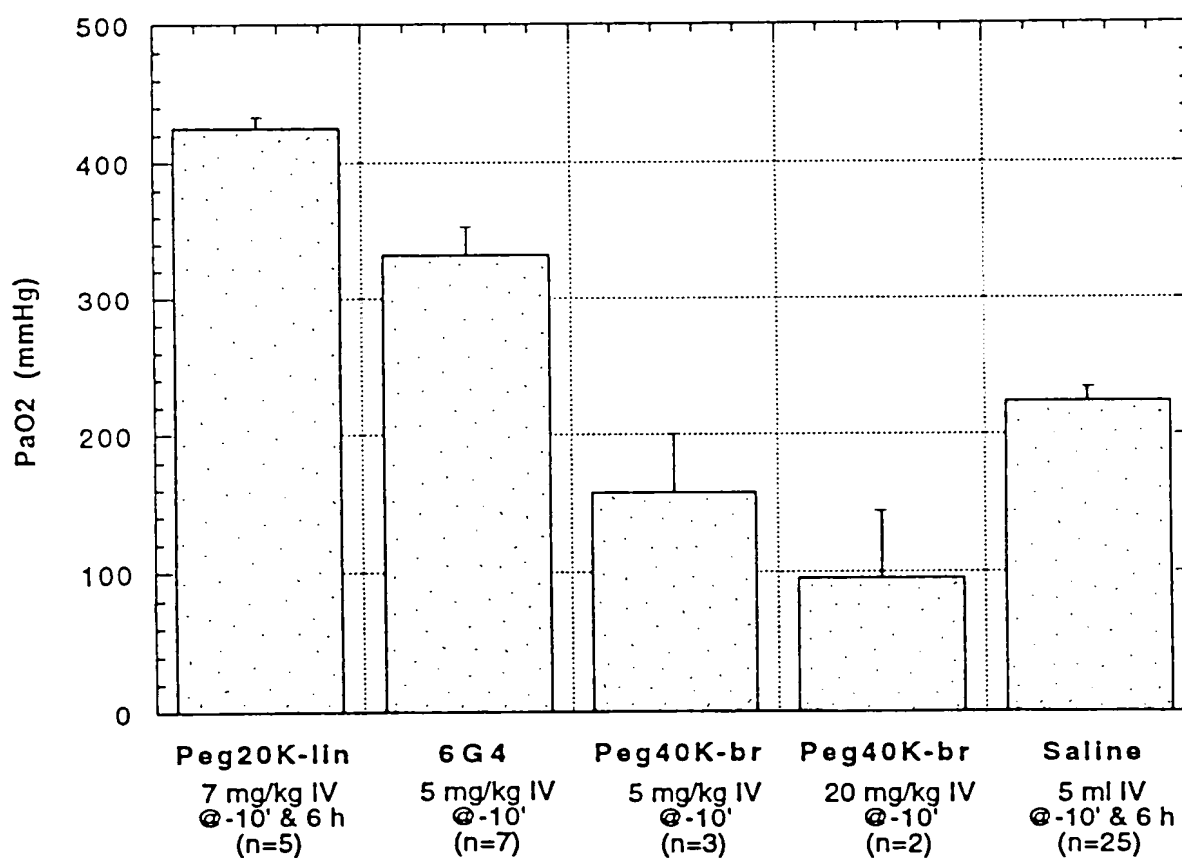


FIG. 70A

JAN 7 4 2003

Oxygenation in 100% O₂ @48 h post Anti-IL8 Tx

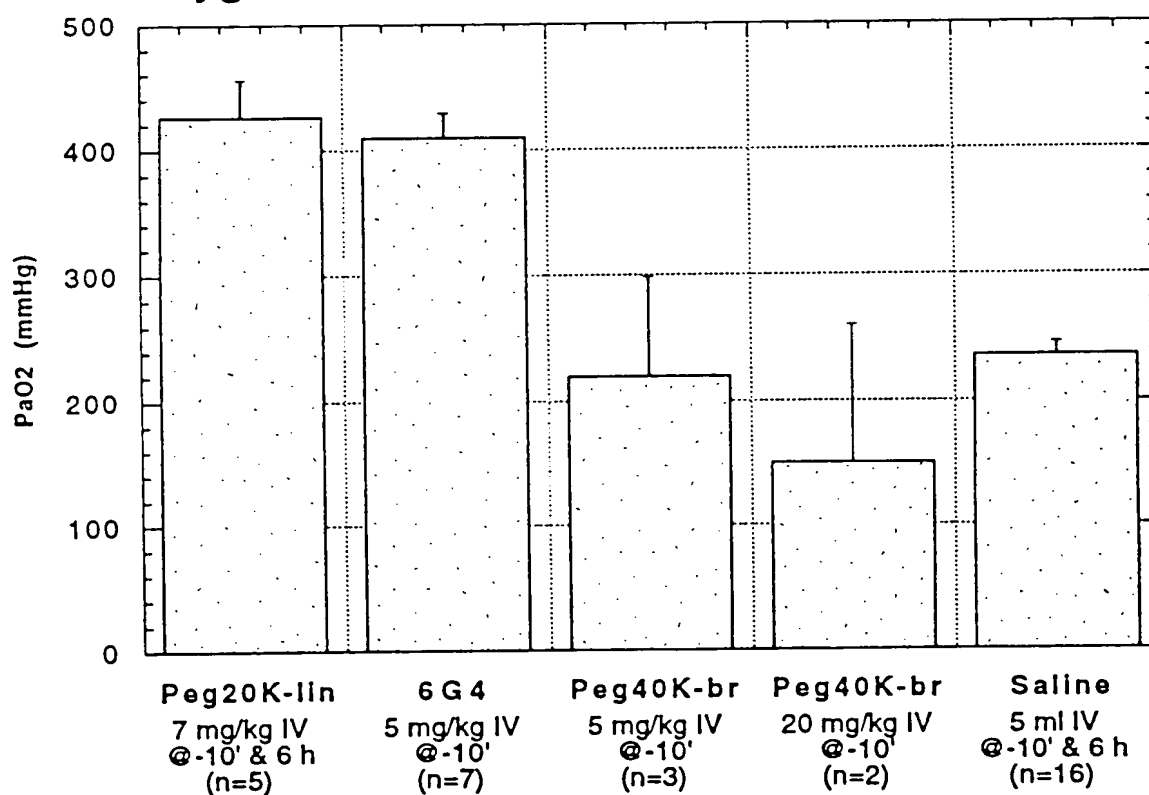


FIG. 70B

JAN 22 2003

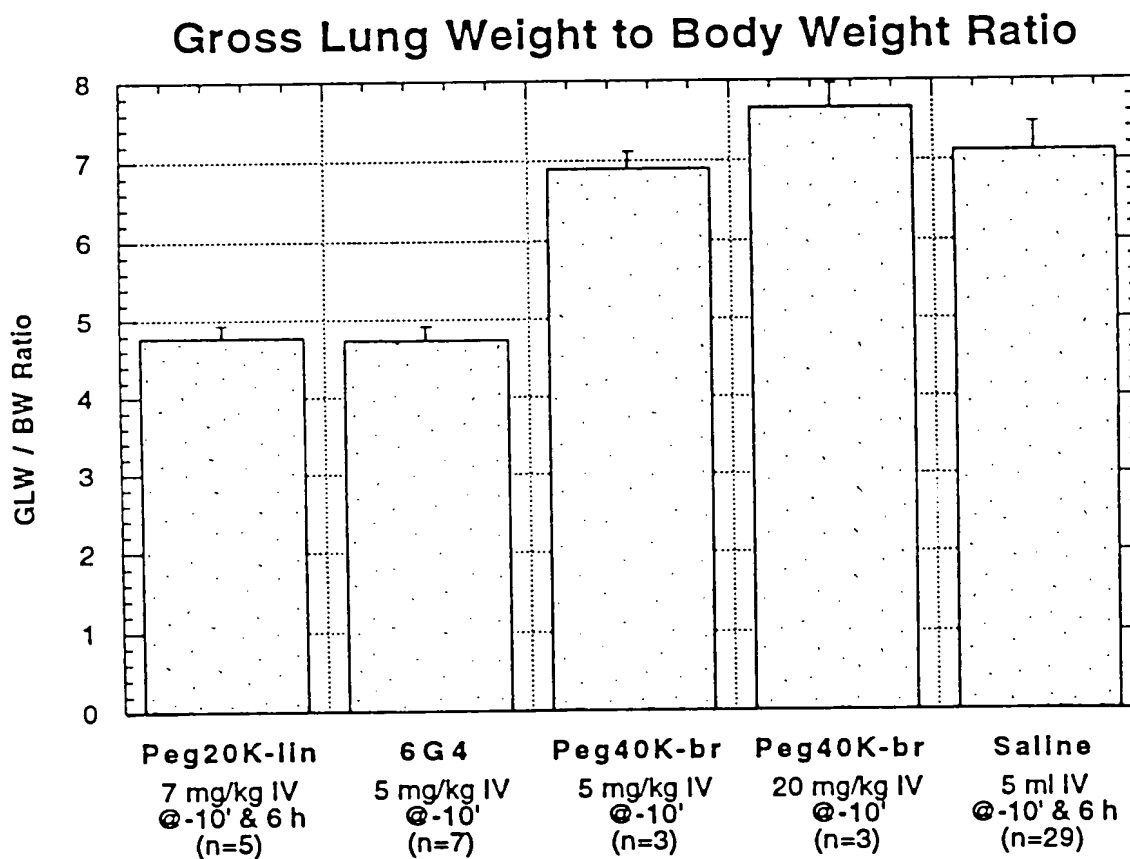


FIG. 70C

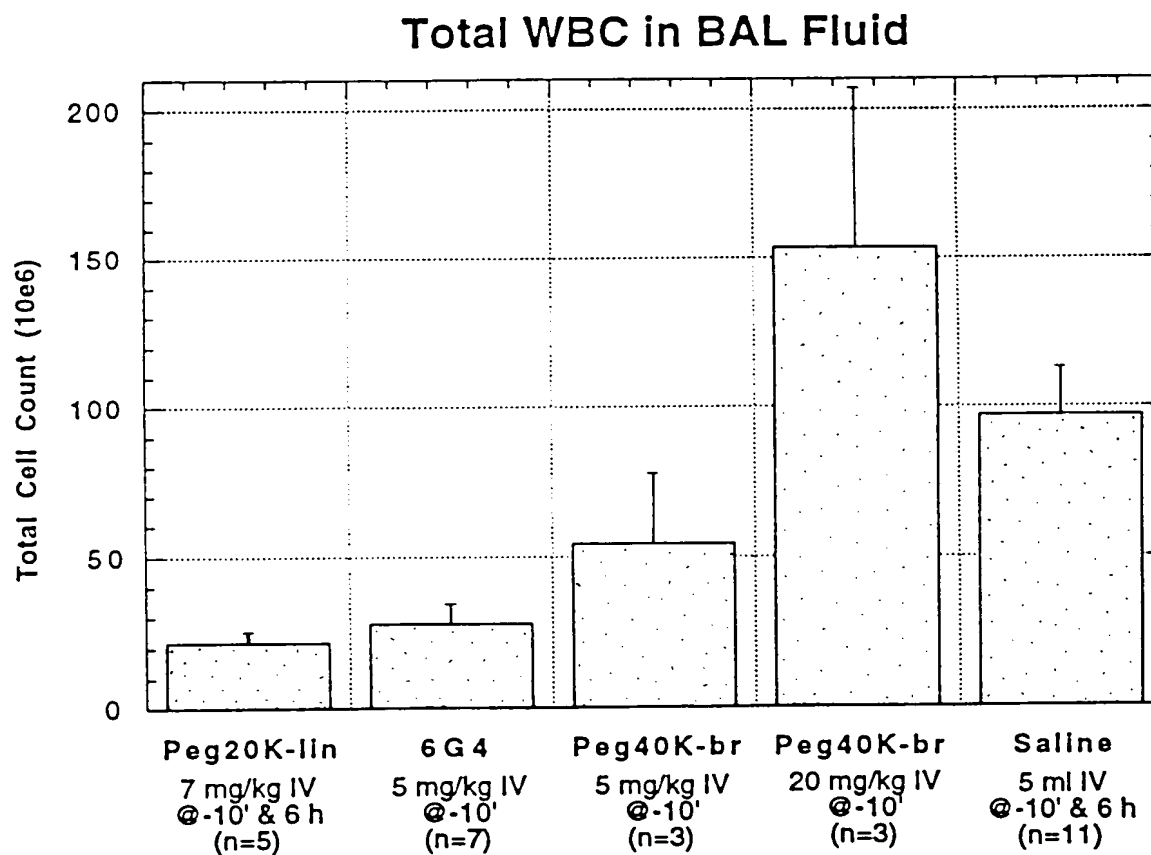
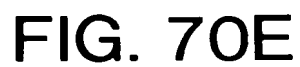
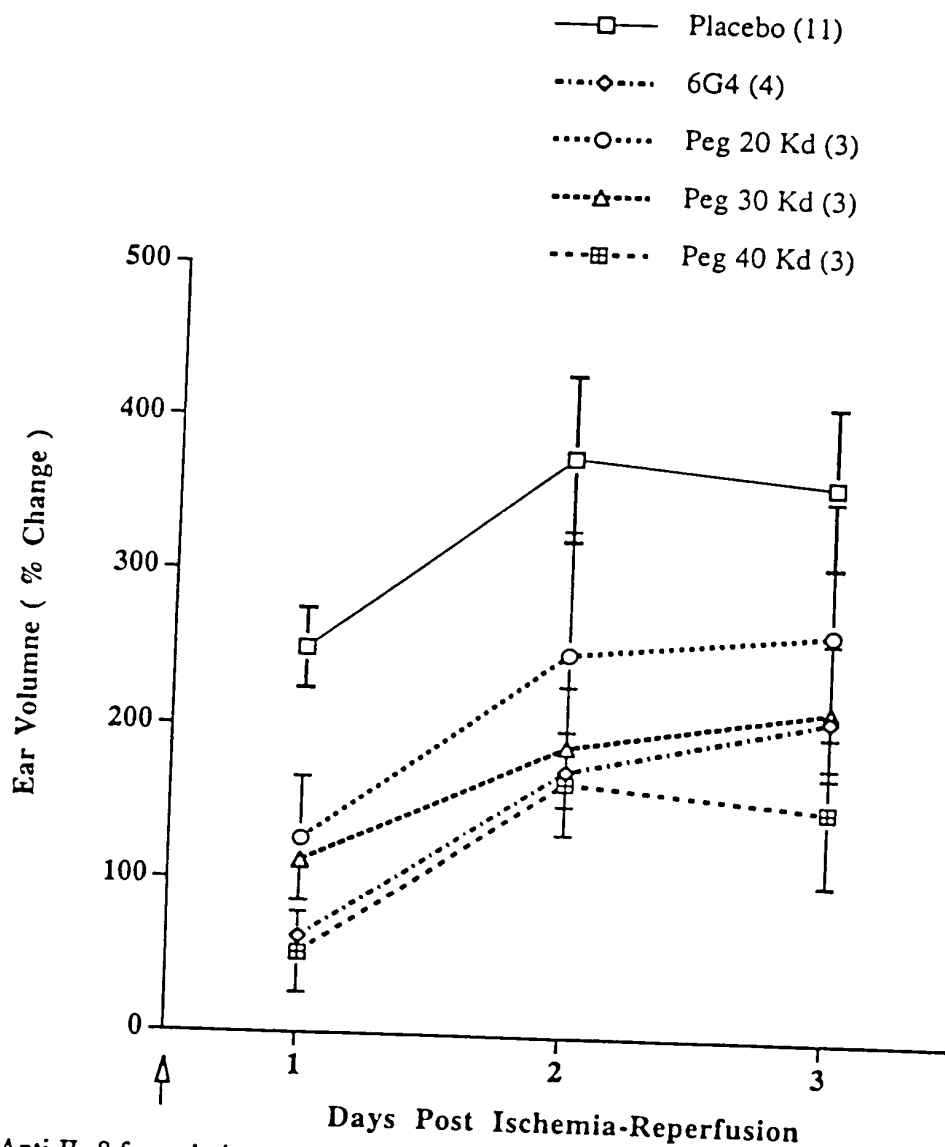


FIG. 70D





The Effect of Pegylated Anti-IL-8 in the Rabbit Ear model of Ischemia-Reperfusion Injury



Anti-IL-8 formulations :
Single Dose (5 mg/kg)
administered IV at time
of reperfusion

FIG. 71